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(54) Title: HUMAN PROTEINS HAVING TRANSMEM	BRAN	E DOMAINS AND DNAS ENCODING THESE PROTEINS		
(57) Abstract				
Proteins comprising any of the amino acid sequences of the nucelotide sequences of SEQ ID NOS: 19 to 36 are		Q ID NOS: 1 to 18 and DNAs encoding said proteins and comprising any ed.		
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#### DESCRIPTION

# Human Proteins Having Transmembrane Domains and DNAs Encoding These Proteins

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#### FIELD OF THE INVENTION

The present invention relates to human proteins having transmembrane domains and cDNAs encoding these proteins. The membrane proteins of this invention can be used as pharmaceuticals or as antigens for preparing antibodies against said proteins. The cDNAs of the invention can be used as probes for the gene diagnosis and gene sources for the gene therapy. The cDNAs can also be used as gene sources for large-scale production of the membrane proteins encoded by the same. The cells into which the genes encoding the membrane proteins are introduced for expression of such membrane proteins in large amounts can be used for detection of the corresponding ligands, screening of low molecular weight medicines, etc.

#### 20 BACKGROUND OF THE INVENTION

Membrane proteins play important roles as signal receptors, ion channels, transporters, etc. for the material transportation or information transmission mediated by the cell membrane. For instance, they are known to serve as receptors for various cytokines, ion channels for sodium ion, potassium ion, chloride ion, etc., transporters for saccharides and amino acids, and so on. The genes for many of them have been cloned already.

In recent years, it was clarified that the abnormalities

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of these membrane proteins are related to a number of hitherto cryptogenic diseases. For example, a gene for a membrane protein having 12 transmembrane domains was identified as the gene responsible for cystic fibrosis [Rommens, J. M. et al., 5 Science 245: 1059-1065 (1989)]. It was also clarified that several membrane proteins act as the receptors when a virus infects the cells. For example, HIV-1 was revealed to infect into the cells through the mediation of a membrane protein fusin, a membrane protein on the T-cell membrane, having a CD-4 antigen and 7 transmembrane domains [Feng, Y. et al., Science 272: 872-877 (1996)]. Therefore, the discovery of new membrane proteins is anticipated to lead to the elucidation of the causes of many diseases, and the isolation of new genes coding for the membrane proteins is desired.

Heretofore, owing to the difficulty in their purification, many of membrane proteins have been isolated by an approach from the gene side. A general method is the so-called expression cloning which comprises transfection of a cDNA library in the animal cells to express the cDNA and detection of the cells expressing the target membrane protein on the membrane by an immunological technique using an antibody or a physiological technique for the change in the membrane permeability. However, this method is applicable only to cloning of a gene for a membrane protein with a known function.

In general, membrane proteins possess hydrophobic transmembrane domains inside the proteins which are synthesized in
the ribosome. Said domains remain in the phospholipid to be
trapped in the membrane. Accordingly, the evidence of the cDNA
for encoding the membrane protein is provided by determination

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of the whole base sequence of a full-length cDNA and detection of highly hydrophobic transmembrane domains in the amino acid sequence of the protein encoded by said cDNA.

As a result of the extensive study, there have successful
ly been obtained human proteins having transmembrane domains,
particularly comprising any of the amino acid sequences of SEQ

ID NOS: 1 to 18, by cloning cDNAs coding for proteins having
transmembrane domains, particularly comprising any of the
nucleotide sequences of SEQ ID NOS: 19 to 36, from a human

full-length cDNA bank. The present invention is based on the
above success.

#### SUMMARY OF THE INVENTION

A main object of the present invention is to provide novel

human proteins having transmembrane domains, particularly
comprising any of the amino acid sequences of SEQ ID NOS: 1 to

18. Another object of this invention is to provide DNAs coding
for said novel proteins, particularly comprising any of the
nucleotide sequences of SEQ ID NOS: 19 to 36. A further object

of the invention is to provide expression vectors capable of in
vitro translating said DNAs or expressing said DNAs in
eukaryotic cells. A still further object of the invention is
to provide transformed eukaryotic cells capable of expressing
said DNAs to produce said proteins.

In one embodiment, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of the amino acid sequences of SEQ ID NOS: 1 to 18 and their fragments.

In another embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of the nucleotide sequences of SEQ ID NOS: 19 to 36.

In a further embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of the nucleotide sequences of SEQ ID NOS: 37 to 54.

#### 10 BRIEF DESCRIPTION OF DRAWINGS

- Figure 1: A figure depicting the structure of the secretory signal sequence detection vector pSSD3.
- Figure 2: A figure depicting the hydrophobicity/hydrophilicity profile of the protein encoded by clone HP01263.
- 15 Figure 3: A figure depicting the hydrophobicity/hydrophilicity profile of the protein encoded by clone HP01299.
  - Figure 4: A figure depicting the hydrophobicity/hydrophilicity profile of the protein encoded by clone HP01347.
- Figure 5: A figure depicting the hydrophobicity/hydrophi-20 licity profile of the protein encoded by clone HP01440.
  - Figure 6: A figure depicting the hydrophobicity/hydrophilicity profile of the protein encoded by clone HP01526.
  - Figure 7: A figure depicting the hydrophobicity/hydrophilicity profile of the protein encoded by clone HP10230.
- 25 Figure 8: A figure depicting the hydrophobicity/hydrophilicity profile of the protein encoded by clone HP10389.
  - Figure 9: A figure depicting the hydrophobicity/hydrophilicity profile of the protein encoded by clone HP10408.
    - Figure 10: A figure depicting the hydrophobicity/hydro-

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philicity profile of the protein encoded by clone HP10412.

Figure 11: A figure depicting the hydrophobicity/hydrophilicity profile of the protein encoded by clone HP10413.

Figure 12: A figure depicting the hydrophobicity/hydrophilicity profile of the protein encoded by clone HP10415.

Figure 13: A figure depicting the hydrophobicity/hydrophilicity profile of the protein encoded by clone HP10419.

Figure 14: A figure depicting the hydrophobicity/hydrophilicity profile of the protein encoded by clone HP10424.

10 Figure 15: A figure depicting the hydrophobicity/hydrophilicity profile of the protein encoded by clone HP10428.

Figure 16: A figure depicting the hydrophobicity/hydrophilicity profile of the protein encoded by clone HP10429.

Figure 17: A figure depicting the hydrophobicity/hydro15 philicity profile of the protein encoded by clone HP10432.

Figure 18: A figure depicting the hydrophobicity/hydrophilicity profile of the protein encoded by clone HP10433.

Figure 19: A figure depicting the hydrophobicity/hydrophilicity profile of the protein encoded by clone HP10480.

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#### BEST MODE FOR CARRING OUT INVENTION

The proteins of the present invention can be obtained, for example, by isolation from human organs, cell lines, etc., by chemical synthesis on the basis of the amino acid sequences as herein disclosed, or by recombinant DNA technology using the DNA encoding the transmembrane domains of the invention. Among them, adoption of the recombinant DNA technology is preferred. Specifically, each of the proteins may be prepared by in vitro transcription of a vector comprising the cDNA of the invention

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to make RNA and in vitro translation using this RNA as a template to accomplish in vitro expression. Also, each of the proteins may be prepared in a large amount by the use of Escherichia coli, Bacillus subtilis, yeasts, animal cells, etc. 5 comprising a suitable expression vector having the DNA encoding such protein.

In the case of producing the protein of the invention by the use of a microorganism such as Escherichia coli, the translation region of the cDNA of the invention is constructed in an expression vector having an origin, a promoter, a ribosome-binding site, a cDNA-cloning site, a terminator, etc. that can be replicated in the microorganism and, after transformation of the host cells with said expression vector, the resultant transformant is incubated, whereby the protein 15 encoded by said cDNA can be produced in a large amount in the microorganism. In that case, a protein fragment containing an optional region can be obtained by performing the expression with inserting an initiation codon and a termination codon before and after the optional translation region. Alternative-20 ly, a fusion protein with another protein can be expressed. Only a protein portion encoding said cDNA can be obtained by cleavage of said fusion protein with an appropriate protease.

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For production of the protein of the invention by expression of DNA coding for such protein in eukaryotic cells, 25 the translation region of said cDNA may be recombined into an expression vector for eukaryotic cells having a promoter, a splicing domain, a poly(A) addition site, etc., followed by introduction into eukaryotic cells so that the protein of the invention is produced as a membrane protein on the cell

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membrane surface. Examples of the expression vector are pKA1, pED6\_dpc2, pCDM8, pSVK3, pMSG, pSVL, pBK-CMV, pBK-RSV, EBV vector, pRS, pYES2, etc. As the eukaryotic cells, there are exemplified mammalian animal culture cells (e.g. simian kidney cells COS7, chinese hamster ovary cells CHO), budding yeasts, Schizosaccharomyces pombe, silkworm cells, Xenopus laevis egg cells, etc., but any other eukaryotic cells may also be used insofar as the protein of the invention can be expressed on the membrane surface. In order to introduce the expression vector into eukaryotic cells, there may be adopted any conventional procedure such as electroporation, calcium phosphate method, liposome method or DEAE dextran method.

The proteins of the present invention include peptide fragments (5 or more amino acid residues) containing any 15 partial amino acid sequence of the amino acid sequences of SEQ ID NOS: 1 to 18. These fragments can be used as antigens for preparation of the antibodies. Also, the proteins of the invention that have signal sequences appear in the form of maturation proteins on the cell surface, after the signal 20 sequences are removed. Therefore, these maturation proteins shall come within the scope of the present invention. The Nterminal amino acid sequences of the maturation proteins can be easily identified by using the method for the cleavage-site determination in a signal sequence [Japan Patent Kokai No. 25 187100/96]. Further, many membrane proteins are subjected to the processing on the cell surface to be converted to the secretor forms. These secretor proteins or peptides shall come within the scope of the present invention. When glycosylation sites are present in the amino acid sequences, expression in

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appropriate animal cells affords glycosylated proteins. Therefore, these glycosylated proteins or peptides also shall come within the scope of the invention.

The DNAs of the invention include all DNAs encoding the above-mentioned proteins. Said DNAs can be obtained using the method by chemical synthesis, the method by cDNA cloning, and so on.

Each of the cDNAs of the invention can be cloned from, for example, the cDNA libraries of the human cell origin. The cDNA is synthesized using as a template a poly(A)<sup>+</sup> RNA extracted from human cells. The human cells may be cells delivered from the human body, for example, by the operation or may be the culture cells. The cDNA can be synthesized by using any method selected from the Okayama-Berg method [Okayama, H. and Berg, P., Mol. Cell. Biol. 2: 161-170 (1982)], the Gubler-Hoffman method [Gubler, U. and Hoffman, J. Gene 25: 263-269 (1983)], and so on, but it is preferred to use the capping method [Kato, S. et al., Gene 150: 243-250 (1994)] as illustrated in Examples in order to obtain a full-length clone in an effective manner.

The primary selection of a cDNA encoding a human protein having transmembrane domains is performed by the sequencing of a partial base sequence of the cDNA clone selected at random from the cDNA libraries, sequencing of the amino acid sequence encoded by the base sequence, and recognition of the presence or absence of hydrophobic site(s) in the resulting N-terminal amino acid sequence region. Next, the secondary selection is carried out by determination of the whole base sequence by the sequencing and the protein expression by the in vitro translation. The ascertainment of the cDNA of the present

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invention for encoding the protein having the secretory signal sequence is performed by using the signal sequence detection method [Yokoyama-Kobayashi, M. et al., Gene 163: 193-196 (1995)]. In other words, the ascertainment for the coding portion of the inserted cDNA fragment to function as a signal sequence is provided by fusing a cDNA fragment encoding the N-terminus of the target protein with a cDNA encoding the protease domain of urokinase and then expressing the resulting cDNA in COS7 cells to detect the urokinase activity in the cell culture medium. On the other hand, the N-terminal region is judged to remain in the membrane in the case where the urokinase activity is not detected in the cell culture medium.

The cDNAs of the invention are characterized by containing any of the nucleotide sequences of SEQ ID NOS: 19 to 36 or any of the nucleotide sequences of SEQ ID NOS: 37 to 54. Table 1 summarizes the clone number (HP number), the cells affording the cDNA, the total nucleotide number of the cDNA, and the number of the amino acid residues of the encoded protein, for each of the cDNAs.

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Table 1

5	Sequence Number	HP Number	Cells	Number of Nucleotides	Number of Amino Acid Residues
10	1, 19, 37	HP01263	Liver	1502	382
	2, 20, 38	HP01299	Liver	1349	317
	3, 21, 39	HP01347	Liver	1643	296
15	4, 22, 40	HP01440	Stomach cancer	729	197
	5, 23, 41	HP01526	Stomach cancer	1322	221
20	6, 24, 42	HP10230	Stomach cancer	3045	251
20	7, 25, 43	HP10389	КВ	653	106
	8, 26, 44	HP10408	Stomach cancer	439	78
25	9, 27, 45	HP10412	Stomach cancer	1131	314
	10, 28, 46	HP10413	Stomach cancer	1875	195
30	11, 29, 47	HP10415	Stomach cancer	1563	462
30	12, 30, 48	HP10419	Stomach cancer	2030	247
	13, 31, 49	HP10424	Stomach cancer	493	113
35	14, 32, 50	HP10428	KB .	2044	365
	15, 33, 51	HP10429	Stomach cancer	1043	226
40	16, 34, 52	HP10432	Liver	972	129
-10	17, 35, 53	HP10433	Liver	695	163
	18, 36, 54	HP10480	Stomach cancer	1914	193

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Hereupon, the same clone as any of the cDNAs of the invention can be easily obtained by screening of the cDNA libraries constructed from the cell line or the human tissues employed in the invention, by the use of an oligonucleotide probe synthesized on the basis of the corresponding cDNA nucleotide sequence of SEQ ID NOS: 37 to 54.

In general, the polymorphism due to the individual difference is frequently observed in human genes. Therefore, any cDNA that is subjected to insertion or deletion of one or plural nucleotides and/or substitution with other nucleotides

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in SEQ ID NOS: 37 to 54 shall come within the scope of the invention.

In a similar manner, any protein that is produced by these modifications comprising insertion or deletion of one or plural nucleotides and/or substitution with other nucleotides shall come within the scope of the present invention, as far as said protein possesses the activity of the corresponding protein having the amino acid sequence of SEQ ID NOS: 1 to 18.

The cDNAs of the invention include cDNA fragments (more than 10 bp) containing any partial nucleotide sequence of the nucleotide sequence of SEQ ID NOS: 19 to 36 or of the nucleotide sequence of SEQ ID NOS: 37 to 54. Also, DNA fragments consisting of a sense chain and an anti-sense chain shall come within this scope. These DNA fragments can be used as the probes for the gene diagnosis.

The present invention also provides genes corresponding to the polynucleotide sequences disclosed herein. "Corresponding genes" are the regions of the genome that are transcribed to produce the mRNAs from which cDNA polynucleotide sequences are derived and may include contiguous regions of the genome necessary for the regulated expression of such genes. Corresponding genes may therefore include but are not limited to coding sequences, 5' and 3' untranslated regions, alternatively spliced exons, introns, promoters, enhancers, and silencer or suppressor elements. The corresponding genes can be isolated in accordance with known methods using the sequence information disclosed herein. Such methods include the preparation of probes or primers from the disclosed sequence information for identification and/or amplification of genes in appropriate

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genomic libraries or other sources of genomic materials. An "isolated gene" is a gene that has been separated from the adjacent coding sequences, if any, present in the genome of the organism from which the gene was isolated.

5 Organisms that have enhanced, reduced, or modified expression of the gene(s) corresponding to the polynucleotide sequences disclosed herein are provided. The desired change in gene expression can be achieved through the use of antisense polynucleotides or ribozymes that bind and/or cleave 10 the mRNA transcribed from the gene (Albert and Morris, 1994, Trends Pharmacol. Sci. 15(7): 250-254; Lavarosky et al., 1997, Biochem. Mol. Med. 62(1): 11-22; and Hampel, 1998, Prog. Nucleic Acid Res. Mol. Biol. 58: 1-39; all of which are incorporated by reference herein). Transgenic animals that 15 have multiple copies of the gene(s) corresponding to the polynucleotide sequences disclosed herein, preferably produced by transformation of cells with genetic constructs that are stably maintained within the transformed cells and their progeny, are provided. Transgenic animals that have modified 20 genetic control regions that increase or reduce gene expression levels, or that change temporal or spatial patterns of gene expression, are also provided (see European Patent No. 0 649 464 Bl, incorporated by reference herein). In addition, organisms are provided in which the gene(s) corresponding to 25 the polynucleotide sequences disclosed herein have been partially or completely inactivated, through insertion of extraneous sequences into the corresponding gene(s) or through deletion of all or part of the corresponding gene(s). Partial or complete gene inactivation can be accomplished through

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insertion, preferably followed by imprecise excision, transposable elements (Plasterk, 1992, Bioessays 14(9): 629-633; Zwaal et al., 1993, Proc. Natl. Acad. Sci. USA 90(16): 7431-7435; Clark et al., 1994, Proc. Natl. Acad. Sci. USA 91(2): 719-722; all of which are incorporated by reference herein), or through homologous recombination, preferably detected by positive/negative genetic selection strategies (Mansour et al., 1988, Nature 336: 348-352; U.S. Patent Nos. 5,464,764; 5,487,992; 5,627,059; 5,631,153; 5,614, 5,616,491; and 5,679,523; all of which are incorporated by reference herein). These organisms with altered gene expression are preferably eukaryotes and more preferably are mammals. Such organisms are useful for the development of non-human models for the study of disorders involving the 15 corresponding gene(s), and for the development of assay systems for the identi fication of molecules that interact with the protein product(s) of the corresponding gene(s).

Where the protein of the present invention membrane-bound (e.g., is a receptor), the present invention 20 also provides for soluble forms of such protein. In such forms part or all of the intracellular and transmembrane domains of the protein are deleted such that the protein is fully secreted from the cell in which it is expressed. The intracellular and transmembrane domains of proteins of the invention can be 25 identified in accordance with known techniques for determination of such domains from sequence information.

Proteins and protein fragments of the present invention include proteins with amino acid sequence lengths that are at least 25% (more preferably at least 50%, and most preferably at

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least 75%) of the length of a disclosed protein and have at least 60% sequence identity (more preferably, at least 75% identity; most preferably at least 90% or 95% identity) with that disclosed protein, where sequence identity is determined 5 by comparing the amino acid sequences of the proteins when aligned so as to maximize overlap and identity while minimizing sequence gaps. Also included in the present invention are proteins and protein fragments that contain a segment preferably comprising 8 or more (more preferably 20 or more, 10 most preferably 30 or more) contiguous amino acids that shares at least 75% sequence identity (more preferably, at least 85% identity; most preferably at least 95% identity) with any such segment of any of the disclosed proteins.

Species homologs of the disclosed polynucleotides and proteins are also provided by the present invention. As used herein, a "species homologue" is a protein or polynucleotide with a different species of origin from that of a given protein or polynucleotide, but with significant sequence similarity to the given protein or polynucleotide, as determined by those of skill in the art. Species homologs may be isolated and identified by making suitable probes or primers from the sequences provided herein and screening a suitable nucleic acid source from the desired species.

The invention also encompasses allelic variants of the 25 disclosed polynucleotides or proteins; that is, naturally-occurring alternative forms of the isolated polynucleotide which also encode proteins which are identical, homologous, or related to that encoded by the polynucleotides.

The invention also includes polynucleotides with sequences

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complementary to those of the polynucleotides disclosed herein.

The present invention also includes polynucleotides capable of hybridizing under reduced stringency conditions, more preferably stringent conditions, and most preferably 5 highly stringent conditions, to polynucleotides described herein. Examples of stringency conditions are shown in the table below: highly stringent conditions are those that are at least as stringent as, for example, conditions A-F; stringent conditions are at least as stringent as, for example, conditions G-L; and reduced stringency conditions are at least as stringent as, for example,

Table 2

Stringency	Polynucleotide	Hybrid	Hybridization Temperature	Wash
Condition	Hybrid	Length	and Buffer <sup>†</sup>	Temperature
		(bp) <sup>‡</sup>		and Buffer <sup>†</sup>
A	DNA : DNA	≥50	65℃; 1×SSC -or-	65°C; 0.3×SSC
			42℃; 1×SSC,50% formamide	
В	DNA : DNA	<50	T <sub>B</sub> *; 1×SSC	T <sub>B</sub> *; 1×SSC
C	DNA : RNA	≥50	67℃; 1×SSC -or-	67°C; 0.3×SSC
			45°C; 1×SSC,50% formamide	
D	DNA: RNA	<50	T <sub>D</sub> *; 1×SSC	T <sub>D</sub> *; 1×SSC
${f E}$	RNA: RNA	≥50	70℃; 1×SSC -or-	70°C; 0.3×SSC
-			50℃; 1×SSC,50% formamide	
F	RNA: RNA	<50	T <sub>F</sub> *; 1×SSC	T <sub>F</sub> *; 1×SSC
G	DNA : DNA	≥50	65°C; 4×SSC -or-	65℃; 1×SSC
			42℃; 4×SSC,50% formamide	
H	DNA : DNA	<50	T <sub>H</sub> *; 4×SSC	T <sub>H</sub> *; 4×SSC
I	DNA: RNA	≥50	67°C; 4×SSC -or-	67°C; 1×SSC
			45°C; 4×SSC,50% formamide	
J	DNA: RNA	<50	T <sub>J</sub> *; 4×SSC	T <sub>J</sub> *; 4×SSC
K	RNA: RNA	≥50	70°C; 4×SSC -or-	67℃; 1×SSC
·			50°C; 4×SSC,50% formamide	
L	RNA: RNA	<50	T <sub>L</sub> *; 2×SSC	T <sub>L</sub> *; 2×SSC
M	DNA : DNA	≥50	50°C; 4×SSC -or-	50°C; 2×SSC
			40℃; 6×SSC,50% formamide	
N	DNA : DNA	<50	T <sub>N</sub> *; 6×SSC	T <sub>N</sub> *; 6×SSC
0	DNA : RNA	≥50	55°C; 4×SSC -or-	55°C; 2×SSC
			42℃; 6×SSC,50% formamide	-
P	DNA : RNA	<50	T <sub>P</sub> *; 6×SSC	T <sub>P</sub> *; 6×SSC
Q	RNA: RNA	≥50	60℃; 4×SSC -or-	60℃; 2×SSC
			45℃; 6×SSC,50% formamide	
R	RNA: RNA	<50	T <sub>R</sub> *; 4×SSC	T <sub>R</sub> *; 4×SSC

- ‡: The hybrid length is that anticipated for the hybridized region(s) of the hybridizing polynucleotides. When hybridizing a polynucleotide to a target polynucleotide of unknown sequence, the hybrid length is assumed to be that of the hybridizing polynucleotide. When polynucleotides of known sequence are hybridized, the hybrid length can be determined by aligning the sequences of the polynucleotides and identifying the region or regions of optimal sequence complementarity.
- †: SSPE (1×SSPE is 0.15M NaCl, 10mM NaH<sub>2</sub>PO<sub>4</sub>, and 1.25mM EDTA, pH7.4) can be substituted for SSC (1×SSC is 0.15M NaCl and 15mM sodium citrate) in the hybridization and wash buffers; washes are performed for 15 minutes after hybridization is complete.
- \* $T_B$   $T_R$ : The hybridization temperature for hybrids anticipated to be less than 50 base pairs in length should be 5-10°C less than the melting temperature ( $T_m$ ) of the hybrid, where  $T_m$  is determined according to the following equations. For hybrids less than 18 base pairs in length,  $T_m$ (°C)=2(#of A + T bases) + 4(# of G + C bases). For hybrids between 18 and 49 base pairs in length,  $T_m$ (°C)=81.5 + 16.6(log<sub>10</sub>[Na<sup>+</sup>]) + 0.41 (%G+C) (600/N), where N is the number of bases in the hybrid, and [Na<sup>+</sup>] is the concentration of sodium ions in the hybridization buffer ([Na<sup>+</sup>] for 1×SSC=0.165M).

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Additional examples of stringency conditions for polynucleotide hybridization are provided in Sambrook, J., E.F. Fritsch, and T. Maniatis, 1989,

Molecular Cloning: A Laboratory

5 Manual, Cold Spring Harbor Laboratory Press, Cold Spring Harbor, NY, chapters 9 and 11, and Current Protocols in Molecular Biology, 1995, F.M. Ausubel et

al., eds., John Wiley & Sons, Inc.,

where sequence identity is

sections 2.10 and 6.3-6.4, incorporated herein by reference.

- 10 Preferably, each such hybridizing polynucleotide has a length that is at least 25%(more preferably at least 50%, and most preferably at least 75%) of the length of the polynucleotide of the present invention to which it hybridizes, and has at least 15 60% sequence identity (more
  - preferably, at least 75% identity; most preferably at least 90% or 95% identity) with the polynucleotide of the present invention to which it hybridizes,
- 20 determined by comparing the sequences of the hybridizing polynucleotides when aligned so as to maximize overlap and identity while minimizing sequence

#### 25 EXAMPLE

gaps.

The present invention is embodied in more detail by the following examples, but this embodiment is not intended to restrict the present invention. The basic operations and the enzyme reactions with regard to the DNA recombination are

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Laboratory Manual", Cold Spring Harbor Laboratory, 1989]. Unless otherwise stated, restrictive enzymes and a variety of modification enzymes to be used were those available from Takara Shuzo Co., Ltd. The manufacturer's instructions were used for the buffer compositions as well as for the reaction conditions, in each of the enzyme reactions. The cDNA synthesis was carried out according to the literature [Kato, S. et al., Gene 150: 243-250 (1994)].

(1) Preparation of Poly(A) + RNA

The epidermoid carcinoma cell line KB (ATCC CRL 17), tissues of stomach cancer delivered by the operation, and liver were used for human cells to extract mRNAs. The cell line was cultured by a conventional procedure.

15 After about 1 g of human tissues was homogenized in 20 ml of a 5.5 M guanidinium thiocyanate solution, total mRNAs were prepared in accordance with the literature [Okayama, H. et al., "Methods in Enzymology" Vol. 164, Academic Press, 1987]. These mRNAs were subjected to chromatography using an oligo(dT)-cellulose column washed with 20 mM Tris-hydrochloric acid buffer solution (pH 7.6), 0.5 M NaCl, and 1 mM EDTA to obtain a poly(A)<sup>+</sup> RNA in accordance with the above-mentioned literature.

#### (2) Construction of cDNA Library

To a solution of 10 μg of the above-mentioned poly(A)<sup>+</sup> RNA in 100 mM Tris-hydrochloric acid buffer solution (pH 8) was added one unit of an RNase-free, bacterium-origin alkaline phosphatase and the resulting solution was allowed to react at 37°C for one hour. After the reaction solution underwent the

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phenol extraction followed by the ethanol precipitation, the obtained pellets were dissolved in a mixed solution of 50 mM sodium acetate (pH 6), 1 mM EDTA, 0.1% 2-mercaptoethanol, and 0.01% Triton X-100. Thereto was added one unit of a tobacco-origin pyrophosphatase (Epicenter Technologies) and the resulting solution at a total volume of 100 µl was allowed to react at 37°C for one hour. After the reaction solution underwent the phenol extraction followed by the ethanol precipitation, the thus-obtained pellets were dissolved in water to obtain a decapped poly(A)<sup>†</sup> RNA solution.

To a solution of the decapped poly(A)<sup>+</sup> RNA and 3 nmol of a DNA-RNA chimeric oligonucleotide (5'-dG-dG-dG-dG-dA-dA-dT-dT-dC-dG-dA-G-G-A-3') in a mixed aqueous solution of 50 mM Trishydrochloric acid buffer solution (pH 7.5), 0.5 mM ATP, 5 mM MgCl<sub>2</sub>, 10 mM 2-mercaptoethanol, and 25% polyethylene glycol were added 50 units of T4 RNA ligase and the resulting solution at a total volume of 30  $\mu$ l was allowed to react at 20°C for 12 hours. After the reaction solution underwent the phenol extraction followed by the ethanol precipitation, the thusobtained pellets were dissolved in water to obtain a chimeric oligo-capped poly(A)<sup>+</sup> RNA.

After the vector pKA1 developed by the present inventors (Japanese Patent Kokai Publication No. 1992-117292) was digested with KpnI, an about 60-dT tail was inserted by a terminal transferase. This product was digested with EcoRV to remove the dT tail at one side and the resulting molecule was used as a vectorial primer.

After 6  $\mu g$  of the previously-prepared chimeric oligocapped poly(A)<sup>+</sup> RNA was annealed with 1.2  $\mu g$  of the vectorial

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primer, the product was dissolved in a mixed solution of 50 mM Tris-hydrochloric acid buffer solution (pH 8.3), 75 mM KCl, 3 mM MgCl<sub>2</sub>, 10 mM dithiothreitol, and 1.25 mM dNTP (dATP + dCTP + dGTP + dTTP), mixed with 200 units of a reverse transferase (GIBCO-BRL), and the resulting solution at a total volume of 20 µl was allowed to react at 42°C for one hour. After the reaction solution underwent the phenol extraction followed by the ethanol precipitation, the thus-obtained pellets were dissolved in a mixed solution of 50 mM Tris-hydrochloric acid 10 buffer solution (pH 7.5), 100 mM NaCl, 10 mM MgCl2, and 1 mM dithiothreitol. Thereto were added 100 units of EcoRI and the resulting solution at a total volume of 20  $\mu$ l was allowed to react at 37°C for one hour. After the reaction solution underwent the phenol extraction followed by the ethanol 15 precipitation, the obtained pellets were dissolved in a mixed solution of 20 mM Tris-hydrochloric acid buffer solution (pH 7.5), 100 mM KCl, 4 mM MgCl<sub>2</sub>, 10 mM (NH<sub>4</sub>)<sub>2</sub>SO<sub>4</sub>, and 50  $\mu$ g/ml bovine serum albumin. Thereto were added 60 units of Escherichia coli DNA ligase and the resulting solution was 20 allowed to react at 16°C for 16 hours. To the reaction solution were added 2 µl of 2 mM dNTP, 4 units of Escherichia coli DNA polymerase I, and 0.1 unit of Escherichia coli DNase H and the resulting solution was allowed to react at 12°C for one hour and then at 22°C for one hour.

Next, the cDNA-synthesis reaction solution was used to transform *Escherichia coli* DH12S (GIBCO-BRL). The transformation was carried out by the electroporation method. A portion of the transformant was inoculated on a 2xYT agar culture medium containing 100 µg/ml ampicillin, which was

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incubated at 37°C overnight. A colony grown on the culture medium was randomly picked up and inoculated on 2 ml of the 2xYT culture medium containing 100  $\mu g/ml$  ampicillin, which was incubated at 37°C overnight. The culture medium was centrifuged 5 to separate the cells, from which a plasmid DNA was prepared by the alkaline lysis method. After the plasmid DNA was doubledigested with EcoRI and NotI, the product was subjected to 0.8% agarose gel electrophoresis to determine the size of the cDNA insert. In addition, by the use of the obtained plasmid as a 10 template, the sequence reaction using M13 universal primer labeled with a fluorescent dye and Taq polymerase (a kit of Applied Biosystems Inc.) was carried out and the product was analyzed by a fluorescent DNA-sequencer (Applied Biosystems Inc.) to determine the base sequence of the cDNA 5'-terminal of 15 about 400 bp. The sequence data were filed as a homo-protein cDNA bank data base.

(3) Selection of cDNAs Encoding Proteins Having
Transmembrane Domains

The base sequence registered in the homo-protein cDNA bank

20 data base was converted to three frames of amino acid sequences
and the presence or absence of an open reading frame (ORF)
beginning from the initiation codon. Then, the selection was
made for the presence of a signal sequence that is
characteristic to a secretory protein at the N-terminal of the

25 portion encoded by ORF. These clones were sequenced from the
both 5' and 3' directions by using the deletion method to
determine the sequence of the whole base sequence. The
hydrophobicity/hydrophilicity profiles were obtained for
proteins encoded by ORF by the Kyte-Doolittle method [Kyte, J.

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& Doolittle, R. F., J. Mol. Bio. 157: 105-132 (1982)] to examine the presence or absence of a hydrophobic region. In the case in which there is a hydrophobic region of putative transmembrane domain(s) in the amino acid sequence of an encoded protein, this protein was considered as a membrane protein.

(4) Construction of Secretory Signal Detection Vector pSSD3

One microgram of pSSD1 carrying the SV40 promoter and a 10 cDNA encoding the protease domain of urokinase [Yokoyama-Kobayashi, M. et al., Gene 163: 193-196 (1995)] was digested with 5 units of Bg1II and 5 units of EcoRV. Then, after dephosphorylation at the 5' terminal by the CIP treatment, a DNA fragment of about 4.2 kbp was purified by cutting off from the gel of agarose gel electrophoresis.

Two oligo DNA linkers, L1 (5'-GATCCCGGGTCACGTGGGAT-3') and L2 (5'-ATCCCACGTGACCCGG-3'), were synthesized phosphorylated by T4 polynucleotide kinase. After annealing of the both linkers, followed by ligation with the previously-20 prepared pSSD1 fragment by T4 DNA ligase, Escherichia coli JM109 was transformed. A plasmid pSSD3 was prepared from the transformant and the objective recombinant was confirmed by the determination of the base sequence of the linker-inserted fragment. Figure 1 illustrates the structure of the thus-25 obtained plasmid. The present plasmid vector carries three types of blunt-end formation restriction enzyme sites, SmaI, PmaCI, and EcoRV. Since these cleavage sites are positioned in succession at an interval of 7 bp, selection of an appropriate site in combination of three types of frames for the inserting

cDNA allows to construct a vector expressing a fusion protein.

(5) Functional Verification of Secretory Signal Sequence Whether the N-terminal hydrophobic region in the secretory protein clone candidate obtained in the above-mentioned steps functions as the secretory signal sequence was verified by the method described in the literature [Yokoyama-Kobayashi, M. et al., Gene 163: 193-196 (1995)]. First, the plasmid containing the target cDNA was cleaved at an appropriate restriction enzyme site that existed at the downstream of the portion 10 expected for encoding the secretory signal sequence. In the case in which this restriction enzyme site was a protruding terminus, the site was blunt-ended by the Klenow treatment or treatment with the mung-bean nuclease. Digestion with HindIII was further carried out and a DNA fragment containing the SV40 15 promoter and a cDNA encoding the secretory sequence at the downstream of the promoter was separated by agarose gel electrophoresis. This fragment was inserted between the pSSD3 HindIII site and a restriction enzyme site selected so as to match with the urokinase-coding frame, thereby constructing a 20 vector expressing a fusion protein of the secretory signal portion of the target cDNA and the urokinase protease domain.

After Escherichia coli (host: JM109) bearing the fusionprotein expression vector was incubated at 37°C for 2 hours in
2 ml of the 2xYT culture medium containing 100 μg/ml
25 ampicillin, the helper phage M13KO7 (50 μl) was added and the
incubation was continued at 37°C overnight. A supernatant
separated by centrifugation underwent precipitation with
polyethylene glycol to obtain single-stranded phage particles.
These particles were suspended in 100 μl of 1 mM Tris-0.1 mM

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EDTA, pH 8 (TE). Also, there was used as a control a suspension of single-stranded particles prepared in the same manner from the vector pLA1-UPA containing pSSD3 and a full-length cDNA of urokinase [Yokoyama-Kobayashi, M. et al., Gene 163: 193-196 (1995)].

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The simian-kidney-origin culture cells, COS7, incubated at 37°C in the presence of 5% CO2 in the Dulbecco's modified Eagle's culture medium (DMEM) containing 10% bovine fetus albumin. Into a 6-well plate (Nunc Inc., 3 cm in the well 10 diameter) were inoculated  $1 \times 10^5$  COS7 cells and incubation was carried out at 37°C for 22 hours in the presence of 5% CO2. After the culture medium was removed, the cell surface was washed with a phosphate buffer solution and then washed again with DMEM containing 50 mM Tris-hydrochloric acid (pH 7.5) (TDMEM). To the cells were added 1  $\mu$ l of the single-stranded 15 phage suspension, 0.6 ml of the DMEM culture medium, and 3 µl of TRANSFECTAM<sup>TM</sup> (IBF Inc.) and the resulting mixture was incubated at 37°C for 3 hours in the presence of 5% CO2. After the sample solution was removed, the cell surface was washed 20 with TDMEM, 2 ml per well of DMEM containing 10% bovine fetus albumin was added, and the incubation was carried out at 37°C for 2 days in the presence of 5% CO<sub>2</sub>.

To 10 ml of 50 mM phosphate buffer solution (pH 7.4) containing 2% bovine fibrinogen (Miles Inc.), 0.5% agarose, and 1 mM potassium chloride were added 10 units of human thrombin (Mochida Pharmaceutical Co., Ltd.) and the resulting mixture was solidified in a plate of 9 cm in diameter to prepare a fibrin plate. Ten microliters of the culture supernatant of the

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transfected COS7 cells were spotted on the fibrin plate, which was incubated at 37°C for 15 hours. The diameter of the thusobtained clear circle was taken as an index for the urokinase activity. In the case in which a cDNA fragment codes for the 
amino acid sequence that functions as a secretory signal sequence, a fusion protein is secreted to form a clear circle 
by its urokinase activity. Therefore, in the case in which a 
clear circle is not formed, the fusion protein remains as 
trapped in the membrane and the cDNA fragment is considered to 
code for a transmembrane domain.

#### (6) Protein Synthesis by In Vitro Translation

The plasmid vector carrying the cDNA of the present invention was utilized for the transcription/translation by the  $T_NT$  rabbit reticulocyte lysate kit (Promega Biotec). In this 15 case, [35]methionine was added and the expression product was labeled with the radioisotope. All reactions were carried out by following the protocols attached to the kit. Two micrograms of the plasmid was allowed to react at 30°C for 90 minutes in total 25 ml of a reaction solution containing 12.5 µl of the 20  $T_NT$  rabbit reticulocyte lysate, 0.5  $\mu$ l of the buffer solution (attached to the kit), 2 µl of an amino acid mixture (methionine-free), 2  $\mu$ l (0.37 MBq/ $\mu$ l) of [ $^{35}$ S]methionine (Amersham Corporation), 0.5 µl of T7 RNA polymerase, and 20 U of RNasin. To 3  $\mu$ l of the reaction solution was added 2  $\mu$ l of an SDS sampling buffer (125 mM Tris-hydrochloric acid suffer solution, pH 6.8, 120 mM 2-mercaptoethanol, 2% SDS solution, 0.025% bromophenol blue, and 20% glycerol) and the resulting solution was heated at 95°C for 3 minutes and then subjected to SDS-polyacrylamide gel electrophoresis. The molecular weight of

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the translation product was determined by carrying out the autoradiography.

#### (7) Expression in COS7

Escherichia coli bearing a vector expressing the protein

of the invention was infected with helper phage M13KO7, and single-stranded phage particles were obtained according to the method as stated above. Using the thus obtained phages, each expression vecotr was introduced into simian-kidney-origin culture cells COS7 in the manner as stated above. After incubation at 37 °C for 2 days in the presence of 5 % CO<sub>2</sub>, further incubation was carried out in a medium containing [35S]cysteine or [35S]methionine for 1 hour. The cells were collected, dissolved and then subjected to SDS-PAGE whereby a band corresponding to the expression product of each protein which is not present in COS7 cells was revealed. In Table 3, the molecular weight of each expression product is shown.

Table 3

HP Number	Supernatant of culture	Membrane fraction
450-504	(kDa)	(kDa)
HP01263	50	-
HP01299	-	30
HP01526	-	22
HP10230	- -	24
HP10408		7
HP10415	-	45
HP10424	<b>-</b>	14
HP10429	-	27
HP10432	-	17
HP10480	_	22

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#### (8) Clone Examples

<HP01263> (Sequence Number 1, 19, 37)

Determination of the whole base sequence for the cDNA insert of clone HP01263 obtained from the human liver cDNA 5 libraries revealed the structure consisting of a 5'-nontranslation region of 36 bp, an ORF of 1149 bp, and a 3'-nontranslation region of 316 bp. The ORF codes for a protein consisting of 382 amino acid residues with one transmembrane domain at the N-terminal. Figure 2 depicts the hydrophobicity 10 /hydrophilicity profile of the present protein obtained by the The in vitro translation resulted in Kyte-Doolittle method. formation of a translation product of 42 kDa, which is almost consistent with the molecular weight of 42,054 as predicted On expression in COS cells, an expression from the ORF. 15 product of about 50 kDa was observed in the culture supernatant. Therefore, said protein can be understood to be a secreted protein. Application of the rule (-3, -1) as a method for anticipation of a cutting site in a secretion signal sequence suggested that the mature protein would start from methionine at 19 position.

The search of the protein data base using the amino acid sequence of the present protein revealed that the protein was analogous to the human  $\alpha$ -2-HS-glycoprotein (SWISS-PROT Accession No. P02765). Table 4 indicates the comparison of the amino acid sequences between the human protein of the present invention (HP) and the human  $\alpha$ -2-HS-glycoprotein (GP). - represents a gap, \* represents an amino acid residue identical to that in the protein of the present invention, and . represents an amino acid residue analogous to that in the

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protein of the present invention. The both proteins possessed a homology of 25.5%. The cysteine position is reserved and this region is analogous to that in cystatins (thiol proteinase inhibitors). There are observed other analogy with histidine-rich glycoprotein (P04196, 30.9%/194 amino acid residues), kininogen (P01045, 24.1%/261 amino acid residues), tyrosine kinase inhibitor (A32827, 24.4%/291 amino acid residues), and so on.

#### Table 4

10 HP MGLLLPLALCILVLCCGAMSPPQLALNPSALLSR--GCNDSDVLAVAGFALRDINKDRKD .\*.\*\* ... . \* . ..\*. \* .\*.\*... ..\* \*. \*\*.. MKSLVLLLCLAQLWGCHSAPHGPGLIYRQPNCDDPETEEAALVAIDYINQNLPW GP HP GYVLRLNRVNDAQEYRRGGLGSLFYLTLDVLETDCHVLRKKAWQDCGMRIFFE-SVYGQC 15 GP GYKHTLNQIDEVKVWPQQPSGELFEIEIDTLETTCHVLDPTPVARCSVRQLKEHAVEGDC HP K-AIFYMNNPSRVLYLAAYNCTLRPVSKKKIYMTCPDCPSSIPTDSSNHQVLEAATESLA GP DFQLLKLDGKFSVVY---AKCDSSPDSAEDVRKVCQDCPLLAPLN--DTRVVHAAKAALA 20 HP KYNNENTSKQYSLFKVTRASSQWVVGPSYFVEYLIKESPC---TKSQASSCSLQSSDSVP . \*\* .\*\*. .... \* . ..\*..\* ... .\*..\*... \* ...\*\* GP AFNAQNNGSNFQLEEISRAQLV-PLPPSTYVEFTVSGTDCVAKEATEAAKCNLLAEKQY-HP VGLCKGSLTRTHWEKFVSVTCDFFESQAPATGSENSAVNQK-PTNLPKVEESQQKNTPPT \*.\*\*..\*. . \*.\*\*\*. \*..\*. .. ..... \*\* . . .... 25 GP -GFCKATLSEKLGGAEVAVTCTVFQTQPVTSQPQPEGANEAVPTPVVDPDAPPSPPLGAP HP DSPSKAGPRGSVQYLPDLDDKNSQEKGPQEAFPVHLDLTTNPQGETLDISFLFLEPMEEK . \*. ..\*..\* \*. GP GLPPAGSPPDSHVLLAAPPGHQLHRAHYDLRHTFMGVVSLGSPSGEVSHPRKTRTVVQPS HP LVVLPFPKEKARTAECPGPAQNASPLVLPP 30 GP VGAAAGPVVPPCPGRIRHFKV

Furthermore, the search of GenBank using the base sequence of the present cDNA revealed that there existed some ESTs possessing the homology of 90% or more (for example, Accession No. H57204), but it can not be assessed whether these ESTs with partial sequences code for the same protein as the protein of the present invention. Hereupon, most of ESTs matching with the present cDNA are available from liver cDNA libraries, whereby the present clone is considered to be expressed specifically in the liver.

The present protein, because of being a type-II membrane protein, is considered to exert its function as a receptor on the membrane surface with the C-terminal side exposed outside the cells or after undergoing a processing followed by being excreted in the serum. The present protein, because of bearing a cystatin-like domain, is considered to possess a proteinase-inhibitor activity as well as many physiological activities in the same manner as for other members of this family. In addition, the present protein, because of being expressed specifically in liver cells, is considered to play a significant role for maintaining the liver function.

<HP01299> (Sequence Number 2, 20, 38)

Determination of the whole base sequence for the cDNA insert of clone HP01299 obtained from the human liver cDNA libraries revealed the structure consisting of a 5'-non-translation region of 110 bp, an ORF of 954 bp, and a 3'-non-translation region of 285 bp. The ORF codes for a protein consisting of 317 amino acid residues with two or more transmembrane domains. Figure 3 depicts the hydrophobicity/hydrophilicity profile of the present protein

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obtained by the Kyte-Doolittle method. The in vitro translation resulted in the formation of a translation product of 32 kDa that was almost consistent with the molecular weight of 35,965 predicted from the ORF.

The search of the protein data base using the amino acid sequence of the present protein revealed that the protein was analogous to the rat retinol dehydrogenase (NBRF Accession No. A55884). Table 5 indicates the comparison of the amino acid sequences between the human protein of the present invention (HP) and the rat retinol dehydrogenase (RN). - represents a gap, \* represents an amino acid residue identical to that in the protein of the present invention, and represents an amino acid residue analogous to that in the protein of the present invention. The both proteins possessed a homology of 65.3% among the entire regions.

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#### Table 5

HP MWLYLAAFVGLYYLLHWYRERQVVSHLQDKYVFITGCDSGFGNLLARQLDARGLRVLAAC 5 RN MWLYLLALVGLWNLLRLFRERKVVSHLQDKYVFITGCDSGFGNLLARQLDRRGMRVLAAC HP LTEKGAEQLRGQTSDRLETVTLDVTKMESIAAATQWVKEHVGDRGLWGLVNNAGILTPIT \*\*\* \* LTEKGAEQLRSKTSDRLETVILDVTKTESIVAATQWVKERVGNRGLWGLVNNAGISVPVG LCEWLNTEDSMNMLKVNLIGVIQVTLSMLPLVRRARGRIVNVSSILGRVAFFVGGYCVSK 10 RN PNEWMRKKDFASVLDVNLLGVIEVTLNMLPLVRKARGRVVNIASTMGRMSLVGGGYCISK HP YGVEAFSDILRREIQHFGVKISIVEPGYFRTGMTNMTQSLERMKQSWKEAPKHIKETYGQ 15 RN YGVEAFSDSLRRELTYFGVKVAIIEPGGFKTNVTNMERLSDNLKKLWDQTTEEVKEIYGE HP QYFDALYNIMKEGLLNCSTNLNLVTDCMEHALTSVHPRTRYSAGWDAKFFFIPLSYLPTS RN KFQDSYMKAMESLVNTCSGDLSLVTDCMEHALTSCHPRTRYSPGWDAKFFYLPMSYLPTF HP LADYILTRSWPKPAQAV 20 \*.\* .. .. \*\*\*.\*. RN LSDAVIHWGSVKPARAL

Furthermore, the search of GenBank using the base sequence of the present cDNA revealed that there existed some ESTs possessing the homology of 90% or more (for example, Accession No. R35197), but any of them was shorter than the present cDNA and did not contain the initiation codon.

The rat retinol dehydrogenase has been found as a 30 microsomal membrane protein participating in the retinoic acid

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biosynthesis in the liver [Chai, X. et al., J. Biol. Chem. 270: 28408-28412 (1995)]. Accordingly, its homologue, the protein of the present invention, is considered to possess a similar function and can be utilized for diagnosis and treatment of diseases caused by the abnormality of this protein.

<HP01347> (Sequence Number 3, 21, 39)

Determination of the whole base sequence for the cDNA insert of clone HP01347 obtained from the human liver cDNA libraries revealed the structure consisting of a 5'-non-10 translation region of 24 bp, an ORF of 891 bp, and a 3'-nontranslation region of 728 bp. The ORF codes for a protein consisting of 296 amino acid residues with one transmembrane N-terminal. 4 domain at the Figure depicts hydrophobicity/hydrophilicity profile of the present protein 15 obtained by the Kyte-Doolittle method. It was indicated that the present protein remained in the membrane from the observation that the urokinase secretion was not identified and the urokinase activity was detected on the membrane surface, upon transduction into the COS7 cells of an expression vector 20 in which a HindIII-SacI fragment (treated with the mung-bean nuclease) containing a cDNA fragment encoding the N-terminal 73 amino acid residues in the present protein was inserted at the HindIII-EcoRV site of pSSD3. Therefore, the present protein is considered to be a type-II membrane protein. The in vitro 25 translation resulted in the formation of a translation product of 33 kDa that was almost consistent with the molecular weight of 33,527 predicted from the ORF.

The search of the protein data base using the amino acid sequence of the present protein revealed that the protein was

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analogous to the human HIV envelope glycoprotein gp120-binding C-type lectin (GenBank Accession No. M98457). Table 6 indicates the comparison of the amino acid sequences between the human protein of the present invention (HP) and the human HIV envelope glycoprotein gp120-binding C-type lectin (CL). represents a gap, \* represents an amino acid residue identical to that in the protein of the present invention, and . represents an amino acid residue analogous to that in the protein of the present invention. The both proteins possessed a homology of 85.6% among 284 amino acid residues. There is observed at the downstream of the transmembrane domain a sequence with seven repetition of Ile-Tyr-Gln-Xaa-Leu-Thr-Xaa-Leu-Lys-Ala-Ala-Val-Gly-Glu-Leu-Xaa-Xaa-Xaa-Ser-Lys-Xaa-Gln-Xaa.

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#### Table 6

	HP	MSDSKEPRVQQLGLLGCLGHGALVLQLLSFMLLAGVLVAI
		******* ***** **** .
5	CL	${\tt MSDSKEPRLQQLGLLEEEQLRGLGFRQTRGYKSLAGCLGHGPLVLQLLSFTLLAGL}$
	HP	${\tt LVQVSKVPSSLSQEQSEQDAIYQNLTQLKAAVGELSEKSKLQEIYQELTQLKAAVGELPE}$
		********** ***** ******************
	CL	${\tt LVQVSKVPSSISQEQSRQDAIYQNLTQLKAAVGELSEKSKLQEIYQELTQLKAAVGELPE}$
	HP	${\tt KSKLQEIYQELTRLKAAVGELPEKSKLQEIYQELTRLKAAVGELPEKSKLQEIYQELTRL}$
10		*********************************
	CL	${\tt KSKLQEIYQELTRLKAAVGELPEKSKLQEIYQELTWLKAAVGELPEKSKMQEIYQELTRL}$
	HP	${\tt KAAVGELPEKSKLQEIYQELTELKAAVGELPEKSKLQEIYQELTQLKAAVGELPDQSKQQ}$
		********* ****** ******* ****** ****** ****
	CL	${\tt KAAVGELPEKSKQQEIYQELTRLKAAVGELPEKSKQQEIYQELTRLKAAVGELPEKSKQQ}$
15	ВP	${\tt QIYQELTDLKTAFERLCRHCPKDWTFFQGNCYFMSNSQRNWHDSVTACQEVRAQLVVIKT}$
		.*****.**.* ****** .****************
	CL	${\tt EIYQELTQLKAAVERLCHPCPWEWTFFQGNCYFMSNSQRNWHDSITACKEVGAQLVVIKS}$
	HP	AEEQLPAVLEQWRTQQ
		**** *. *
20	CL	${\bf AEEQNFLQLQSSRSNRFTWMGLSDLNQEGTWQWVDGSPLLPSFKQYWNRGEPNNVGEEDC}$

Furthermore, the search of GenBank using the base sequence of the present cDNA revealed that there existed some ESTs possessing the homology of 90% or more (for example, Accession No. H90360), but it can not be assessed whether these ESTs with partial sequences code for the same protein as the protein of the present invention.

The present protein, because of being a type-II membrane 30 protein, is considered to exert its function as a receptor on

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the membrane surface with the C-terminal side exposed outside the cells or after undergoing a processing followed by being excreted in the serum. Hereupon, the human HIV envelope glycoprotein gp120-binding C-type lectin that is highly homologous with the present protein has been found as a CD4-independent HIV receptor [Curtis, B. M. et al., Proc. Natl. Acad. Sci. USA 89: 8356-8360 (1992)].

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<HP01440> (Sequence Number 4, 22, 40)

Determination of the whole base sequence for the cDNA

10 insert of clone HP01440 obtained from the human stomach cancer
cDNA libraries revealed the structure consisting of a 5'-nontranslation region of 37 bp, an ORF of 594 bp, and a 3'-nontranslation region of 98 bp. The ORF codes for a protein
consisting of 197 amino acid residues with four transmembrane

15 domains. Figure 5 depicts the hydrophobicity/hydrophilicity
profile of the present protein obtained by the Kyte-Doolittle
method. The in vitro translation resulted in the formation of
a translation product of 21 kDa that was almost consistent with
the molecular weight of 20,822 predicted from the ORF.

The search of the protein data base using the amino acid sequence of the present protein revealed that the protein was analogous to the human tumor-associated antigen L6 (SWISS-PROT Accession No. P30408). Table 7 indicates the comparison of the amino acid sequences between the human protein of the present invention (HP) and the human tumor-associated antigen L6 (L6).

- represents a gap, \* represents an amino acid residue identical to that in the protein of the present invention, and

. represents an amino acid residue analogous to that in the protein of the present invention. The both proteins possessed

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a homology of 47.0% among the entire regions.

#### Table 7

Furthermore, the search of GenBank using the base sequence of the present cDNA revealed that there existed some ESTs possessing the homology of 90% or more and also containing the initiation codon (for example, Accession No. T55097), but many sequences were not distinct and the same ORF as that in the present cDNA was not identified.

25 The human tumor-associated antigen L6 is a member of a membrane antigen TM4 superfamily proteins which are expressed in large quantities on the surface of human tumor cells [Marken, J. S. et al., Proc. Natl. Acad. Sci. USA 89: 3503-3507 (1992)]. Since these membrane antigens are expressed 30 specifically on some specified cells or cancer cells,

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antibodies against these antigens, if constructed, are useful for a variety of diagnoses and as carriers for the drug delivery. In addition, the cells in which genes of these membrane antigens are transduced and the membrane antigens are expressed are applicable for detection of the corresponding ligands and so on.

<HP01526> (Sequence Number 5, 23, 41)

Determination of the whole base sequence for the cDNA insert of clone HP01526 obtained from the human stomach cancer cDNA libraries revealed the structure consisting of a 5'-non-translation region of 83 bp, an ORF of 666 bp, and a 3'-non-translation region of 573 bp. The ORF codes for a protein consisting of 221 amino acid residues with a hydrophobic region of putative six transmembrane domains. Figure 6 depicts the hydrophobicity/hydrophilicity profile of the present protein obtained by the Kyte-Doolittle method. The in vitro translation resulted in the formation of a translation product of 23 kDa that was almost consistent with the molecular weight of 25,030 predicted from the ORF.

The search of the protein data base using the amino acid sequence of the present protein revealed that the protein was analogous to the mouse interstitial cell protein (GenBank Accession No. X96618). Table 8 indicates the comparison of the amino acid sequences between the human protein of the present invention (HP) and the mouse interstitial cell protein (MM). - represents a gap, \* represents an amino acid residue identical to that in the protein of the present invention, and . represents an amino acid residue analogous to that in the protein of the present invention. The both proteins possessed

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a homology of 79.6% among the entire regions.

#### Table 8

Furthermore, the search of GenBank using the base sequence of the present cDNA revealed that there existed some ESTs possessing the homology of 90% or more and also containing the initiation codon (for example, Accession No. H02682), but many sequences were not distinct and the same ORF as that in the present cDNA was not identified.

The mouse interstitial cell protein has been cloned as a membrane protein that is expressed with highly increasing in interstitial cells stimulated by a cytokine [Tagoh, H. et al., Biochem. Biophys. Res. Commun. 221: 744-749 (1996)]. Since these membrane proteins are expressed specifically on some specified cells and cancer cells, antibodies against these

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proteins, if constructed, are useful for a variety of diagnoses and as carriers for the drug delivery. In addition, the cells in which genes of these membrane antigens are transduced and the membrane antigens are expressed are applicable for detection of the corresponding ligands and so on.

<HP10230> (Sequence Number 6, 24, 42)

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Determination of the whole base sequence for the cDNA insert of clone HP10230 obtained from the human stomach cancer cDNA libraries revealed the structure consisting of a 5'-nontranslation region of 190 bp, an ORF of 756 bp, and a 3'-nontranslation region of 2099 bp. The ORF codes for a protein consisting of 251 amino acid residues with at least one domain. transmembrane Figure 7 depicts the hydrophobicity/hydrophilicity profile of the present protein obtained by the Kyte-Doolittle method. The in vitro translation resulted in the formation of a translation product of 30 kDa that was almost consistent with the molecular weight of 28,800 predicted from the ORF.

The search of the protein data base using the amino acid sequence of the present protein revealed that the protein was analogous to the nematode hypothetical protein F25D7.1 (GenBank Accession No. Z78418). Table 9 indicates the comparison of the amino acid sequences between the human protein of the present invention (HP) and the nematode hypothetical protein F25D7.1 (CE). - represents a gap, \* represents an amino acid residue identical to that in the protein of the present invention, and represents an amino acid residue analogous to that in the protein of the present invention. The both proteins possessed a homology of 49.8% among the entire regions.

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#### Table 9

	HS	MSDIGDWFRSIPAITRYWFAATVAVPLVGKLGLISPAYLFL-WPEAFLYRFQIWRPITAT
		* ** .**** ***.*.*. *** *
5	CE	MDLENFLLGIPIVTRYWFLASTIIPLLGRFGFINVQWMFLQW-DLVVNKFQFWRPLTAL
	HS	FYFPVGPGTGFLYLVNLYFLYQYSTRLETGAFDGRPADYLFMLLFNW-ICIVITGLAMDM
		.*.**.* *** .*. ****.**. ** **.******
	CE	IYYPVTPQTGFHWLMMCYFLYNYSKALESETYRGRSADYLFMLIFNWFFCSGLC-MALDI
	HS	QLLMIPLIMSVLYVWAQLNRDMIVSFWFGTRFKACYLPWVILGFNYIIGGSVINELIGNL
10		.*. ****** *.*.* ****** ** * *****. *** * *.
	CE	YFLLEPMVISVLYVWCQVNKDTIVSFWFGMRFPARYLPWVLWGFNAVLRGGGTNELVGIL
	HS	VGHLYFFLMFRYPMDLGGRNFLSTPQFLYRWLPSRRGGVSGFGVPPASMRRAADQNGGGG
		*** *** *
	CE	VGHAYFFVALKYPDEYGV-DLISTPEFLHRLIPDEDGGIHGQDGNIRGARQQPRG
15	HS	RHNWGQGFRLGDQ
		* * * * **
	CE	-HQWPGGVGARLGGN

20 Furthermore, the search of GenBank using the base sequence of the present cDNA revealed that there existed some ESTs possessing the homology of 90% or more and also containing the initiation codon (for example, Accession No. W01493), but many sequences were not distinct and the same ORF as that in the present cDNA was not identified.

<HP10389> (Sequence Number 7, 25, 43)

Determination of the whole base sequence for the cDNA insert of clone HP10389 obtained from the human epidermoid carcinoma cell line KBc cDNA libraries revealed the structure consisting of a 5'-non-translation region of 62 bp, an ORF of

321 bp, and a 3'-non-translation region of 270 bp. The ORF codes for a protein consisting of 106 amino acid residues with a hydrophobic region of putative two transmembrane domains. Figure 8 depicts the hydrophobicity/hydrophilicity profile of the present protein obtained by the Kyte-Doolittle method. The in vitro translation resulted in the formation of a translation product of 12 kDa that was almost consistent with the molecular weight of 11,528 predicted from the ORF.

The search of the protein data base using the amino acid sequence of the present protein revealed that the protein was not analogous to any of known proteins. Furthermore, the search of GenBank using the base sequence of the present cDNA revealed that there existed some ESTs possessing the homology of 90% or more (for example, Accession No. H70816), but many sequences were not distinct and the same ORF as that in the present cDNA was not identified.

<HP10408> (Sequence Number 8, 26, 44)

Determination of the whole base sequence for the cDNA insert of clone HP10408 obtained from the human stomach cancer cDNA libraries revealed the structure consisting of a 5'-non-translation region of 74 bp, an ORF of 237 bp, and a 3'-non-translation region of 128 bp. The ORF codes for a protein consisting of 78 amino acid residues with a putative signal sequence at the N-terminal as well as a sequence of one putative interior transmembrane domain. Figure 9 depicts the hydrophobicity/hydrophilicity profile of the present protein obtained by the Kyte-Doolittle method. It was indicated that the present protein remained in the membrane from the observation that the urokinase secretion was not identified

upon transduction into the COS7 cells of an expression vector in which a HindIII-BglII fragment (after the Klenow treatment) containing a cDNA fragment encoding the N-terminal 70 amino acid residues in the present protein was inserted at the 5 HindIII-EcoRV site of pSSD3. The in vitro translation resulted in the formation of a translation product of 9 kDa that was almost consistent with the molecular weight of 8,396 predicted from the ORF.

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Furthermore, the search of GenBank using the base sequence of the present cDNA revealed that there existed some ESTs possessing the homology of 90% or more (for example, Accession No. T94049), but they were shorter than the present cDNA and molecule containing the initiation codon was not identified.

<HP10412> (Sequence Number 9, 27, 45) 15

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Determination of the whole base sequence for the cDNA insert of clone HP10412 obtained from the human stomach cancer cDNA libraries revealed the structure consisting of a 5'-nontranslation region of 55 bp, an ORF of 945 bp, and a 3'-non-20 translation region of 131 bp. The ORF codes for a protein consisting of 314 amino acid residues with one transmembrane depicts the 10 N-terminal. Figure domain at the hydrophobicity/hydrophilicity profile of the present protein obtained by the Kyte-Doolittle method. It was indicated that the present protein remained in the membrane from the observation that the urokinase secretion was not identified upon transduction into the COS7 cells of an expression vector in which a HindIII-ApaI fragment (treated with mung-bean nuclease) containing a cDNA fragment encoding the N-terminal 65

43

amino acid residues in the present protein was inserted at the HindIII-EcoRV site of pSSD3. The in vitro translation resulted in the formation of a translation product of 44 kDa that was somewhat larger than the molecular weight of 35,610 predicted from the ORF.

The search of the protein data base using the amino acid sequence of the present protein revealed that the protein was analogous to the nematode hypothetical protein of 28.5 kDa (SWISS-PROT Accession No. P34623). Table 10 indicates the comparison of the amino acid sequences between the human protein of the present invention (HP) and the nematode hypothetical protein of 28.5 kDa (CE). - represents a gap, \* represents an amino acid residue identical to that in the protein of the present invention, and . represents an amino acid residue analogous to that in the protein of the present invention. The both proteins possessed a homology of 42.8% in the C-terminal region of 243 amino acid residues.

#### Table 10

HP MVAPVWYLVAAALLVGFILFLTRSRGRAASAGQEPLHNEELAGAGRVAQPGPLEPEEPRA GGRPRRRDLGSRLQAQRRAQRVAWAEA--DENEEEAVILAQEEEGVEKPAETHLSGKIG 5 CE MRRNARRRVNRDEO EDGFVNHMMNDGEDVEDLDGGAEO FEYDEDGKKIG HP AKKLRKLEEKQARKAQREAEEAEREERKRLESQREAEWKKEEERLRLEEEQKEEEE--RK .\* \*\*..\*.... \*\* \* \*\*\*\*\*\* \*..\* \* \*..\*\*\* . \*...\*. \*\* CE KRKAAKLQAKEEKRQMREYEVREREERKRREEER--EKKRDEERAKEEADEKAEEERLRK HP AREEQAQREHEEYLKLKEAFVVEEEGVGETMTEEQSQSFLTEFINYIKQSKVVLLEDLAS CE EREEKERKEHEEYLAMKASFAIEEEG-TDAIEGEEAENLIRDFVDYVKTNKVVNIDELSS HP QVGLRTQDTINRIQDLLAEGTITGVIDDRGKFIYITPEELAAVANFIRQRGRVSIAELAQ 15 . \*\*...\*..\*\*.\*\* . \*\*.\*\*\*\*\*\* . \*\*.\*\*\*\*\* \*.\*. CE HFGLKSEDAVNRLQHFIEEGLVQGVMDDRGKFIYISDEEFAAVAKFINQRGRVSIHEIAE HP ASNSLIAWGRESPAQAPA .\*\*.\*\* . \*.\*. CE QSNRLIRLETPSAAE 20

Furthermore, the search of GenBank using the base sequence of the present cDNA revealed that there existed some ESTs possessing the homology of 90% or more (for example, Accession No. T09311), but it can not be assessed whether these ESTs with partial sequences code for the same protein as the protein of the present invention.

<HP10413> (Sequence Number 10, 28, 46)

Determination of the whole base sequence for the cDNA insert of clone HP10413 obtained from the human stomach cancer

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cDNA libraries revealed the structure consisting of a 5'-nontranslation region of 78 bp, an ORF of 588 bp, and a 3'-nontranslation region of 1209 bp. The ORF codes for a protein consisting of 195 amino acid residues with one transmembrane N-terminal. Figure depicts domain the 11 hydrophobicity/hydrophilicity profile of the present protein obtained by the Kyte-Doolittle method. It was indicated that the present protein remained in the membrane from the observation that the urokinase secretion was not identified upon transduction into the COS7 cells of an expression vector in which a HindIII-PmaCI fragment containing a cDNA fragment encoding the N-terminal 65 amino acid residues in the present protein was inserted at the HindIII-PmaCI site of pSSD3. The in vitro translation resulted in the formation of a translation 15 product of 28 kDa that was somewhat larger than the molecular weight of 21,671 predicted from the ORF.

The search of the protein data base using the amino acid sequence of the present protein revealed that the protein was analogous to the swine steroidal membrane-binding protein (GenBank Accession No. X99714). 20 Table 11 indicates comparison of the amino acid sequences between the human protein of the present invention (HP) and the swine steroidal membrane-binding protein (SS). - represents a gap, \* represents an amino acid residue identical to that in the protein of the 25 present invention, and . represents an amino acid residue analogous to that in the protein of the present invention. The both proteins possessed a homology of 96.4% among the entire regions.

#### Table 11

	HP	${\tt MAAEDVVATGADPSDLESGGLLHEIFTSPLNLLLLGLCIFLLYKIVRGDQPAASGDSDDD}$
		************
5	SS	${\tt MAAEDVAATGADPSELEGGGLLHEIFTSPLNLLLLGLCIFLLYKIVRGDQPAAS-DSDDD}$
	нР	EPPPLPRLKRRDFTPAELRRFDGVQDPRILMAINGKVFDVTKGRKFYGPEGPYGVFAGRD
		**************
	SS	EPPPLPRLKRRDFTPAELRRFDGVQDPRILMAINGKVFDVTKGRKFYGPEGPYGVFAGRD
•	HP	ASRGLATFCLDKEALKDEYDDLSDLTAAQQETLSDWESQFTFKYHHVGKLLKEGEEPTVY
LO		**************
	SS	ASRGLATFCLDKEALKDEYDDLSDLTPAQQETLNDWDSQFTFKYHHVGKLLKEGEEPTVY
	ĦР	SDEEEPKDESARKND
	-	******
	SS	SDEEEPKDESARKND
1 =		

Furthermore, the search of GenBank using the base sequence of the present cDNA revealed that there existed some ESTs possessing the homology of 90% or more (for example, Accession No. AA021062), but many sequences were not distinct and the same ORF as that in the present cDNA was not identified.

<HP10415> (Sequence Number 11, 29, 47)

Determination of the whole base sequence for the cDNA insert of clone HP10415 obtained from the human stomach cancer cDNA libraries revealed the structure consisting of a 5'-non-translation region of 71 bp, an ORF of 1389 bp, and a 3'-non-translation region of 103 bp. The ORF codes for a protein consisting of 462 amino acid residues with one transmembrane domain at the N-terminal. Figure 12 depicts the hydrophobicity/hydrophilicity profile of the present protein

obtained by the Kyte-Doolittle method. The in vitro translation resulted in the formation of a translation product of 48 kDa that was somewhat smaller than the molecular weight of 52,458 predicted from the ORF.

5 The search of the protein data base using the amino acid sequence of the present protein revealed that the protein was analogous to the cytochrome P450 as exemplified by the simian cytochrome P450IIIA8 (SWISS-PROT Accession No. P33268). Table 12 indicates the comparison of the amino acid sequences between 10 the human protein of the present invention (HP) and the simian cytochrome P450IIIA8 (CP). - represents a gap, \* represents an amino acid residue identical to that in the protein of the present invention, and . represents an amino acid residue analogous to that in the protein of the present invention. The 15 both proteins possessed a homology of 21.3% among the entire regions.

48

# Table 12

	HP	MLDFAIFAVTFLLALVGAVLYLYPASRQAAGIPGITPTEEKDGNLPDIVN-SGSLHEF
		.*******
5	CP	MDLIPDLAVETWLLLAVTLVLLYLYGTHSHGLFKKLGIPGPTPLPLLGNILSYRKGFWTF
	HP	LVNLHERYGPVVSFWFGRRLVVSLGTVDVLKQHINPNKTLDPFETMLK-SLLRYQSGGGS
		** * .*. **. ** * *
	CP	DMECYKKYGKVWGFYDGRQPVLAITDPNMIK-TVLVKECYSVFTNRRPFGPVGFMKNAIS
	HP	VSENHMRKKLYENGVTDSLKSNFALLLKLSEELLDKWLSYPET-QHVPLSQHMLGF
10		**. *
	CP	IAEDEEWKRIRSLLSPTFTSGKLKEMVPIIAKYGDVLVRNLRREAETGKPVTLKDVFGAY
	HP-	AMKSVTQMVMGSTF-EDDQEVIRFQKNHGTVWSEIGKGFLDGSLDKNM
		.** .** *. *. * *.
	ÇP	SMDVITSTSFGVNIDSLNNPQDPFVENTKKLLRFDFLDPFFLSITIFPFIIPILEVLNIS
15	HP	TRKKQYEDALMQ-LESVLRNIIKE-RKGR-NFSQHIFIDSLVQGNLNDQQILEDS
		*
	CP	IFPREVTSFLRKSVKRIKESRLKDTQKHRVDFLQLMIDSQNSKETESHKALSDLELVAQS
	HP	MIFSLASCIITAKLCTWAICFLTTSEEVQKKLYEEINQVF-GNGPVTPEKIEQLRYCQHV
		** .** *. *.* .** *** ** * * . * . *
20	CP	IIFIFAGYETTSSVLSFIIYELATHPDVQQKLQEEIDTVLPNKAPPTYDTVLQMEYLDMV
	HP	LCETVRTAKLTPVSAQLQDIEGKIDRFIIPRETLVLYALGVVLQDPNTWPSPHKFDPDRF
		. **.*
	HP	VNETLRIFPIAMRLERVCKKDVEINGIFIPKGVVVMIPSYALHHDPKYWPEPEKFLPERF
	HP	DDELVMKTFSSLGFSGTQECPELRFAYMVTTVLLSVLVKRLHLLSVEGQVIETKYE
25		.** ****** * * *
	CP	SKKNNDNIDPYIYTPFG-SGPRNCIGMRFALMNMKLAIIRVLQNFSFKPCKETQIPLKLR
	HP	LVTSSREEAWITVSKRY
		*
	CP	LGGLLQTEKPIVLKIESRDGTVSGA

Furthermore, the search of GenBank using the base sequence of the present cDNA revealed that there existed some ESTs

possessing the homology of 90% or more (for example, Accession No. AA381169), but it can not be assessed whether these ESTs with partial sequences code for the same protein as the protein of the present invention.

The cytochrome P450 participates in the drug metabolism and can be utilized as a catalyst in organic synthesis reactions such as oxidation and so on.

<HP10419> (Sequence Number 12, 30, 48)

Determination of the whole base sequence for the cDNA insert of clone HP10419 obtained from the human stomach cancer cDNA libraries revealed the structure consisting of a 5'-non-translation region of 170 bp, an ORF of 744 bp, and a 3'-non-translation region of 1116 bp. The ORF codes for a protein consisting of 247 amino acid residues with a hydrophobic region of putative seven transmembrane domains. Figure 13 depicts the hydrophobicity/hydrophilicity profile of the present protein obtained by the Kyte-Doolittle method.

The search of GenBank using the base sequence of the present cDNA revealed that there existed some ESTs possessing the homology of 90% or more (for example, Accession No. AA340663), but it can not be assessed whether these ESTs with partial sequences code for the same protein as the protein of the present invention.

<HP10424> (Sequence Number 13, 31, 49)

Determination of the whole base sequence for the cDNA insert of clone HP10424 obtained from the human stomach cancer cDNA libraries revealed the structure consisting of a 5'-non-translation region of 97 bp, an ORF of 342 bp, and a 3'-non-translation region of 54 bp. The ORF codes for a protein

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consisting of 113 amino acid residues with one transmembrane Figure 14 depicts N-terminal. domain at the hydrophobicity/hydrophilicity profile of the present protein obtained by the Kyte-Doolittle method. It was indicated that the present protein remained in the membrane from the observation that the urokinase secretion was not identified upon transduction into the COS7 cells of an expression vector in which a HindIII-AccI fragment (after the Klenow treatment) containing a cDNA fragment encoding the N-terminal 58 amino 10 acid residues in the present protein was inserted at the HindIII-SmaI site of pSSD3. The in vitro translation resulted in the formation of a translation product of 14 kDa that was somewhat larger than the molecular weight of 12,784 predicted from the ORF.

of the present cDNA revealed that there existed some ESTs possessing the homology of 90% or more (for example, Accession No. AA401979), but it can not be assessed whether these ESTs with partial sequences code for the same protein as the protein of the present invention.

<HP10428> (Sequence Number 14, 32, 50)

Determination of the whole base sequence for the cDNA insert of clone HP10428 obtained from the human epidermoid carcinoma cell line KBc cDNA libraries revealed the structure consisting of a 5'-non-translation region of 287 bp, an ORF of 1098 bp, and a 3'-non-translation region of 659 bp. The ORF codes for a protein consisting of 365 amino acid residues with a hydrophobic region of putative nine transmembrane domains. Figure 15 depicts the hydrophobicity/hydrophilicity profile of

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the present protein obtained by the Kyte-Doolittle method. The result of the in vitro translation did not reveal the formation of distinct bands and only revealed the formation of smeary bands at the high-molecular-weight position.

5 The search of the protein data base using the amino acid sequence of the present protein revealed that the protein was analogous to the baker's yeast hypothetical membrane protein YML038c (NBRF Accession No. S49741). Table 13 indicates the comparison of the amino acid sequences between the human protein of the present invention (HP) and the baker's yeast hypothetical membrane protein YML038c (SC). - represents a gap, \* represents an amino acid residue identical to that in the protein of the present invention, and . represents an amino acid residue analogous to that in the protein of the present invention. The both proteins possessed a homology of 26.3% among the N-terminal region of 281 amino acid residues.

# Table 13

	HP	MGRWALDVAFLWKAVLTLGLVL-LYYCFSIGITFYNKWLTKSFHFPLFMTMLHLA
		**. *.* ***.*
5	sc	MNRTVFLAFVFGWYFCS-IALSIYNRWMFDPKDGLGIGYPVLVTTFHQA
	HP	VIFLFSALSRALVQCSSHRARVVLSWADYLRRVAPTALATALDVGLSNWSFLYVTVS
		*.*. *
	sc	TLWLLSGIYIKLRHKPVKNVLRKNNGFNWSFFLKFLLPTAVASAGDIGLSNVSFQYVPLT
	HP	LYTMTKSSAVLFILIFSLIFKLEELRAALVLVVLLIAGGLFMFTYKSTQ-FN
10		.**** *.*.*. **** ** **
	sc	IYTIIKSSSIAFVLLFGCIFKLEKFHWKLALSVIIMFVGVALMVFKPSDSTSTKNDQALV
	HP	VEGFALVLGASFIGGIRWTLTQMLLQKAELGLQNPIDTMFHLQPLMFLGLFPLFAVFEGL
		. * *******
	sc	IFGSFLVLASSCLSGLRWVYTQLMLRNNPIQTNTAAAVEES-DGALFTENEDNVDNEPVV
15	HP	HLSTSEKIFRFQDT-GLLLRVLGSLFLGGILAFGLGFSEFLLVSRTSSLTLSIAGIFKEV
		.** * * * *
	sc	NLANNKMLENFGESKPHPIHTIHQLAPIMGITLLLTS-LLVEKPFPGIFS-SSIFRLD
	HP	CTLLLAAHLLGDQISLLNWLGFALCLSGISLHVALKALHSRGDGGPKALKGLGSSPDLEL
20	sc	TSNGGVGTETTVLSIVRGIVLLILPGFAVFLLTICEFSILEQTPVLTVSIVGIVKELLTV
	НР	LLRSSQREEGDNEEEEYFVAQGQQ
	sc	IFGIIILSERLSGFYNWLGMLIIMADVCYYNYFRYKQDLLQKYHSVSTQDNRNELKGFQD

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Furthermore, the search of GenBank using the base sequence of the present cDNA revealed that there existed some ESTs possessing the homology of 90% or more (for example, Accession No. AA018345), but it can not be assessed whether these ESTs

with partial sequences code for the same protein as the protein of the present invention.

<HP10429> (Sequence Number 15, 33, 51)

Determination of the whole base sequence for the cDNA insert of clone HP10429 obtained from the human stomach cancer cDNA libraries revealed the structure consisting of a 5'-nontranslation region of 156 bp, an ORF of 681 bp, and a 3'-nontranslation region of 206 bp. The ORF codes for a protein consisting of 226 amino acid residues with four transmembrane 10 domains. Figure 16 depicts the hydrophobicity/hydrophilicity profile of the present protein obtained by the Kyte-Doolittle method. The in vitro translation resulted in the formation of a translation product of 25 kDa that was almost consistent with the molecular weight of 25,321 predicted from the ORF.

The search of the protein data base using the amino acid sequence of the present protein revealed that the protein was not analogous to any known proteins. Furthermore, the search of GenBank using the base sequence of the present cDNA revealed that there existed some ESTs possessing the homology of 90% or 20 more (for example, Accession No. AA315933), but it can not be assessed whether these ESTs with partial sequences code for the same protein as the protein of the present invention.

<HP10432> (Sequence Number 16, 34, 52)

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Determination of the whole base sequence for the cDNA insert of clone HP10429 obtained from the human liver cDNA 25 libraries revealed the structure consisting of a 5'-nontranslation region of 28 bp, an ORF of 390 bp, and a 3'-nontranslation region of 554 bp. The ORF codes for a protein consisting of 129 amino acid residues with a signal-like sequence at the N-terminal and one interior transmembrane domain. Therefore, the present protein is considered to be a type-I membrane protein. Figure 17 depicts the hydrophobicity/hydrophilicity profile of the present protein obtained by the Kyte-Doolittle method.

The search of the protein data base using the amino acid sequence of the present protein revealed that the protein was not analogous to any known proteins. Furthermore, the search of GenBank using the base sequence of the present cDNA revealed that there existed some ESTs possessing the homology of 90% or more (for example, Accession No. T74424), but the same ORF as that in the present cDNA was not identified.

<HP10433> (Sequence Number 17, 35, 53)

Determination of the whole base sequence for the cDNA insert of clone HP10433 obtained from the human liver cDNA libraries revealed the structure consisting of a 5'-nontranslation region of 72 bp, an ORF of 492 bp, and a 3'-nontranslation region of 131 bp. The ORF codes for a protein consisting of 163 amino acid residues with one transmembrane 20 domain the N-terminal. Figure 18 depicts the at hydrophobicity/hydrophilicity profile of the present protein obtained by the Kyte-Doolittle method. It was indicated that the present protein remained in the membrane from the observation that the urokinase secretion was not identified 25 upon transduction into the COS7 cells of an expression vector in which a HindIII-Eco81I fragment (treated with the mung-bean nuclease) containing a cDNA fragment encoding the N-terminal 137 amino acid residues in the present protein was inserted at the HindIII-EcoRV site of pSSD3. Therefore, the present protein

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is considered to be a type-II membrane protein. The in vitro translation resulted in the formation of a translation product of 21 kDa that was almost consistent with the molecular weight of 18,617 predicted from the ORF.

5 The search of the protein data base using the amino acid sequence of the present protein revealed that the protein was not analogous to any known proteins. Furthermore, the search of GenBank using the base sequence of the present cDNA revealed that there existed some ESTs possessing the homology of 90% or 10 more (for example, Accession No. H84693), but many sequences are not distinct and the same ORF as that in the present cDNA was not identified.

<HP10480> (Sequence Number 18, 36, 54)

Determination of the whole base sequence for the cDNA insert of clone HP10480 obtained from the human stomach cancer cDNA libraries revealed the structure consisting of a 5'-non-translation region of 79 bp, an ORF of 582 bp, and a 3'-non-translation region of 1253 bp. The ORF codes for a protein consisting of 193 amino acid residues with four transmembrane domains. Figure 19 depicts the hydrophobicity/hydrophilicity profile of the present protein obtained by the Kyte-Doolittle method. The in vitro translation resulted in the formation of a translation product of 23 kDa that was somewhat larger than the molecular weight of 21,445 predicted from the ORF.

25 The search of the protein data base using the amino acid sequence of the present protein revealed that the protein was not analogous to any known proteins. Furthermore, the search of GenBank using the base sequence of the present cDNA revealed that there existed some ESTs possessing the homology of 90% or

more (for example, Accession No. W93606), but many sequences are not distinct and the same ORF as that in the present cDNA was not identified.

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The present invention provides human proteins having 5 transmembrane domains and cDNAs encoding said proteins. All of the proteins of the present invention are putative proteins controlling the proliferation and differentiation of the cells, because said proteins exist on the cell membrane. Therefore, the proteins of the present invention can be used as 10 pharmaceuticals or as antigens for preparing antibodies against said proteins. Furthermore, said DNAs can be used for the expression of large amounts of said proteins. The cells expressing large amounts of membrane proteins with transfection of these membrane protein genes can be applied to the detection 15 of the corresponding ligands, the screening of novel lowmolecular medicines, and so on.

In addition to the activities and uses described above, the polynucleotides and proteins of the present invention may exhibit one or more of the uses or biological activities (including those associated with assays cited herein) identified below. Uses or activities described for proteins of the present invention may be provided by administration or use of such proteins or by administration or use of polynucleotides encoding such proteins (such as, for example, in gene therapies 25 or vectors suitable for introduction of DNA).

### Research Uses and Utilities

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The polynucleotides provided by the present invention can be used by the research community for various purposes. polynucleotides can be used to express recombinant protein for

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analysis, characterization or therapeutic use; as markers for tissues in which the corresponding protein is preferentially expressed (either constitutively or at a particular stage of tissue differentiation or development or in disease states); as molecular weight markers on Southern gels; as chromosome markers or tags (when labeled) to identify chromosomes or to map related gene positions; to compare with endogenous DNA sequences in patients to identify potential genetic disorders; as probes to hybridize and thus discover novel, related DNA 10 sequences; as a source of information to derive PCR primers for genetic fingerprinting; as a probe to "subtract-out" known the process of discovering other polynucleotides; for selecting and making oligomers for attachment to a "gene chip" or other support, including for 15 examination of expression patterns; to raise anti-protein antibodiesusing DNA immunization techniques; and as an antigen to raise anti-DNA antibodies or elicit another immune response. Where the polynucleotide encodes a protein which binds or potentially binds to another protein (such as, for example, in 20 a receptor-ligand interaction), the polynucleotide can also be used in interaction trap assays (such as, for example, that described in Gyuris et al., Cell 75:791-803 (1993)) to identify polynucleotides encoding the other protein with which binding occurs or to identify inhibitors of the binding interaction.

25 The proteins provided by the present invention can similarly be used in assay to determine biological activity, including in a panel of multiple proteins for high-throughput screening; to raise antibodies or to elicit another immune response; as a reagent (including the labeled reagent) in

assays designed to quantitatively determine levels of the protein (or its receptor) in biological fluids; as markers for tissues in which the corresponding protein is preferentially expressed (either constitutively or at a particular stage of tissue differentiation or development or in a disease state); and, of course, to isolate correlative receptors or ligands. Where the protein binds or potentially binds to another protein (such as, for example, in a receptor-ligand interaction), the protein can be used to identify the other protein with which binding occurs or to identify inhibitors of the binding interaction. Proteins involved in these binding interactions can also be used to screen for peptide or small molecule inhibitors or agonists of the binding interaction.

Any or all of these research utilities are capable of 15 being developed into reagent grade or kit format for commercialization as research products.

Methods for performing the uses listed above are well known to those skilled in the art. References disclosing such methods include without limitation "Molecular Cloning: A 20 Laboratory Manual", 2d ed., Cold Spring Harbor Laboratory Press, Sambrook, J., E.F. Fritsch and T. Maniatis eds., 1989, and "Methods in Enzymology: Guide to Molecular Cloning Techniques", Academic Press, Berger, S.L. and A.R. Kimmel eds., 1987.

### 25 <u>Nutritional Uses</u>

Polynucleotides and proteins of the present invention can also be used as nutritional sources or supplements. Such uses include without limitation use as a protein or amino acid supplement, use as a carbon source, use as a nitrogen source

and use as a source of carbohydrate. In such cases the protein or polynucleotide of the invention can be added to the feed of a particular organism or can be administered as a separate solid or liquid preparation, such as in the form of powder, pills, solutions, suspensions or capsules. In the case of microorganisms, the protein or polynucleotide of the invention can be added to the medium in or on which the microorganism is cultured.

# Cytokine and Cell Proliferation/Differentiation

# 10 Activity

A protein of the present invention may exhibit cytokine, cell proliferation (either inducing or inhibiting) or cell differentiation (either inducing or inhibiting) activity or may induce production of other cytokines in certain cell populations. Many protein factors discovered to date, including all known cytokines, have exhibited activity in one or more factor dependent cell proliferation assays, and hence the assays serve as a convenient confirmation of cytokine activity. The activity of a protein of the present invention is evidenced by any one of a number of routine factor dependent cell proliferation assays for cell lines including, without limitation, 32D, DA2, DA1G, T10, B9, B9/11, BaF3, MC9/G, M+ (preB M+), 2E8, RB5, DA1, 123, T1165, HT2, CTLL2, TF-1, Mo7e and CMK.

25 The activity of a protein of the invention may, among other means, be measured by the following methods:

Assays for T-cell or thymocyte proliferation include without limitation those described in: Current Protocols in Immunology, Ed by J. E. Coligan, A.M. Kruisbeek, D.H.

Margulies, E.M. Shevach, W Strober, Pub. Greene Publishing Associates and Wiley-Interscience (Chapter 3, In Vitro assays for Mouse Lymphocyte Function 3.1-3.19; Chapter 7, Immunologic studies in Humans); Takai et al., J. Immunol. 137:3494-3500, 1986; Bertagnolli et al., J. Immunol. 145:1706-1712, 1990; Bertagnolli et al., Cellular Immunology 133:327-341, 1991; Bertagnolli, et al., J. Immunol. 149:3778-3783, 1992; Bowman et al., J. Immunol. 152: 1756-1761, 1994.

Assays for cytokine production and/or proliferation of spleen cells, lymph node cells or thymocytes include, without limitation, those described in: Po lyclonal T cell stimulation, Kruisbeek, A.M. and Shevach, E.M. In Current Protocols in Immunology. J.E.e.a. Coligan eds. Vol 1 pp. 3.12.1-3.12.14, John Wiley and Sons, Toronto. 1994; and Measurement of mouse and human Interferon γ, Schreiber, R.D. In Current Protocols in Immunology. J.E.e.a. Coligan eds. Vol 1 pp. 6.8.1-6.8.8, John Wiley and Sons, Toronto. 1994.

differentiation proliferation and Assays for include, without hematopoietic and lymphopoietic cells 20 limitation, those described in: Measurement of Human and Murine Interleukin 2 and Interleukin 4, Bottomly, K., Davis, L.S. and Lipsky, P.E. In Current Protocols in Immunology. J.E.e.a. Coligan eds. Vol 1 pp. 6.3.1-6.3.12, John Wiley and Sons, Toronto. 1991; deVries et al., J. Exp. Med. 173:1205-1211, 25 1991; Moreau et al., Nature 336:690-692, 1988; Greenberger et al., Proc. Natl. Acad. Sci. U.S.A. 80:2931-2938, 1983; Measurement of mouse and human interleukin 6 -Nordan, R. In Current Protocols in Immunology. J.E.e.a. Coligan eds. Vol 1 pp. 6.6.1-6.6.5, John Wiley and Sons, Toronto. 1991; Smith et

al., Proc. Natl. Acad. Sci. U.S.A. 83:1857-1861, 1986;
Measurement of human Interleukin 11 - Bennett, F., Giannotti,
J., Clark, S.C. and Turner, K. J. In Current Protocols in
Immunology. J.E.e.a. Coligan eds. Vol 1 pp. 6.15.1 John Wiley
and Sons, Toronto. 1991; Measurement of mouse and human
Interleukin 9 - Ciarletta, A., Giannotti, J., Clark, S.C. and
Turner, K.J. In Current Protocols in Immunology. J.E.e.a.
Coligan eds. Vol 1 pp. 6.13.1, John Wiley and Sons, Toronto.
1991.

10 Assays for T-cell clone responses to antigens (which will identify, among others, proteins that affect APC-T cell interactions as well as direct T-cell effects by measuring proliferation and cytokine production) include, without limitation, those described in: Current Protocols Immunology, Ed by J. E. Coligan, A.M. Kruisbeek, D.H. Margulies, E.M. Shevach, W Strober, Pub. Greene Publishing Associates and Wiley-Interscience (Chapter 3, In Vitro assays for Mouse Lymphocyte Function; Chapter 6, Cytokines and their cellular receptors; Chapter 7, Immunologic studies in Humans); 20 Weinberger et al., Proc. Natl. Acad. Sci. USA 77:6091-6095, 1980; Weinberger et al., Eur. J. Immun. 11:405-411, 1981; Takai et al., J. Immunol. 137:3494-3500, 1986; Takai et al., J. Immunol. 140:508-512, 1988.

### Immune Stimulating or Suppressing Activity

A protein of the present invention may also exhibit immune stimulating or immune suppressing activity, including without limitation the activities for which assays are described herein. A protein may be useful in the treatment of various immune deficiencies and disorders (including severe combined

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immunodeficiency (SCID)), e.g., in regulating (up or down) growth and proliferation of T and/or B lymphocytes, as well as effecting the cytolytic activity of NK cells and other cell populations. These immune deficiencies may be genetic or be caused by viral (e.g., HIV) as well as bacterial orfungal infections, or may result from autoimmune disorders. More specifically, infectious diseases causes by viral, bacterial,

fungal or other infection may be treatable using a protein of the present invention, including infections by HIV, hepatitis

10 viruses, herpesviruses, mycobacteria, Leishmania spp., malaria spp. and various fungal infections such as candidiasis. Of

course, in this regard, a protein of the present invention may

also be useful where a boost to the immune system generally may

be desirable, i.e., in the treatment of cancer.

Autoimmune disorders which may be treated using a protein 15 of the present invention include, for example, connective sclerosis, systemic tissue disease, multiple erythematosus, rheumatoid arthritis, autoimmune pulmonary inflammation, Guillain-Barre syndrome, autoimmune thyroiditis, 20 insulin dependent diabetes mellitis, myasthenia gravis, graft-versus-host disease and autoimmune inflammatory eye disease. Such a protein of the present invention may also to be useful in the treatment of allergic reactions and conditions, such as asthma (particularly allergic asthma) or 25 other respiratory problems. Other conditions, in which immune suppression is desired (including, for example, organ transplantation), may also be treatable using a protein of the present invention.

Using the proteins of the invention it may also be

possible to immune responses, in a number of ways. regulation may be in the form of inhibiting or blocking an immune response already in progress or may involve preventing The functions of the induction of an immune response. 5 activated T cells may be inhibited by suppressing T cell responses or by inducing specific tolerance in T cells, or Immunosuppression of T cell responses is generally an active, non-antigen-specific, process which requires continuous exposure of the T cells to the suppressive agent. Tolerance, 10 which involves inducing non-responsiveness or anergy in T cells, is distinguishable from immunosuppression in that it is generally antigen-specific and persists after exposure to the tolerizing agent has ceased. Operationally, tolerance can be demonstrated by the lack of a T cell response upon reexposure 15 to specific antigen in the absence of the tolerizing agent.

Down regulating or preventing one or more antigen functions (including without limitation B lymphocyte antigen functions (such as , for example, B7)), e.g., preventing high level lymphokine synthesis by activated T cells, will be useful in situations of tissue, skin and organ transplantation and in 20 graft-versus-host disease (GVHD). For example, blockage of T cell function should result in reduced tissue destruction in Typically, in tissue transplants, tissue transplantation. the transplant is initiated through its rejection of 25 recognition as foreign by T cells, followed by an immune reaction that destroys the transplant. The administration of a molecule which inhibits or blocks interaction of a B7 lymphocyte antigen with its natural ligand(s) on immune cells (such as a soluble, monomeric form of a peptide having B7-2

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activity alone or in conjunction with a monomeric form of a

peptide having an activity of another B lymphocyte antigen

(e.g., B7-1, B7-3) or blocking antibody), prior to

transplantation can lead to the binding of the molecule to the

5 natural ligand(s) on the immune cells without transmitting the

corresponding costimulatory signal. Blocking B lymphocyte

antigen function in this matter prevents cytokine synthesis by

immune cells, such as T cells, and thus acts as an

immunosuppressant. Moreover, the lack of costimulation may

also be sufficient to anergize the T cells, thereby inducing

tolerance in a subject. Induction of long-term tolerance by B

lymphocyte antigen-blocking reagents may avoid the necessity of

repeated administration of these blocking reagents. To achieve

sufficient immunosuppression or tolerance in a subject, it may

15 also be necessary to block the function of a combination of B

lymphocyte antigens.

The efficacy of particular blocking reagents in preventing organ transplant rejection or GVHD can be assessed using animal models that are predictive of efficacy in humans. Examples of appropriate systems which can be used include allogeneic cardiac grafts in rats and xenogeneic pancreatic islet cell grafts in mice, both of which have been used to examine the immunosuppressive effects of CTLA4Ig fusion proteins in vivo as described in Lenschow et al., Science 257:789-792 (1992) and Turka et al., Proc. Natl. Acad. Sci USA, 89:11102-11105 (1992). In addition, murine models of GVHD (see Paul ed., Fundamental Immunology, Raven Press, New York, 1989, pp. 846-847) can be used to determine the effect of blocking B lymphocyte antigen function in vivo on the development of that disease.

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Blocking antigen function may also be therapeutically useful for treating autoimmune diseases. Many autoimmune disorders are the result of inappropriate activation of T cells that are reactive against self tissue and which promote the 5 production of cytokines and autoantibodies involved in the pathology of the diseases. Preventing the activation of autoreactive T cells may reduce or eliminate disease symptoms. Administration of reagents which block costimulation of T cells by disrupting receptor: ligand interactions of B lymphocyte 10 antigens can be used to inhibit T cell activation and prevent production of autoantibodies or T cell-derived cytokines which may be involved in the disease process. Additionally, blocking reagents may induce antigen-specific tolerance of autoreactive T cells which could lead to long-term relief from the disease. 15 The efficacy of blocking reagents in preventing or alleviating autoimmune disorders can be determined using a number of well-characterized animal models of human autoimmune diseases. Examples include murine experimental autoimmune encephalitis, systemic lupus erythmatosis in MRL/lpr/lpr mice or NZB hybrid 20 mice, murine autoimmune collagen arthritis, diabetes mellitus in NOD mice and BB rats, and murine experimental myasthenia gravis (see Paul ed., Fundamental Immunology, Raven Press, New York, 1989, pp. 840-856).

Upregulation of an antigen function (preferably a B lymphocyte antigen function), as a means of up regulating immune responses, may also be useful in therapy. Upregulation of immune responses may be in the form of enhancing an existing immune response or eliciting an initial immune response. For example, enhancing an immune response through stimulating B

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lymphocyte antigen function may be useful in cases of viral infection. In addition, systemic viral diseases such as influenza, the commoncold, and encephalitis might be alleviated by the administration of stimulatory forms of B lymphocyte antigens systemically.

Alternatively, anti-viral immune responses may be enhanced in an infected patient by removing T cells from the patient, costimulating the T cells in vitro with viral antigen-pulsed APCs either expressing a peptide of the present invention or together with a stimulatory form of a soluble peptide of the present invention and reintroducing the in vitro activated T cells into the patient. Another method of enhancing anti-viral immune responses would be to isolate infected cells from a patient, transfect them with a nucleic acid encoding a protein of the present invention as described herein such that the cells express all or a portion of the protein on their surface, and reintroduce the transfected cells into the patient. The infected cells would now be capable of delivering a costimulatory signal to, and thereby activate, T cells in vivo.

In another application, up regulation or enhancement of antigen function (preferably B lymphocyte antigen function) may be useful in the induction of tumor immunity. Tumor cells (e.g., sarcoma, melanoma, lymphoma, leukemia, neuroblastoma, carcinoma) transfected with a nucleic acid encoding at least one peptide of the present invention can be administered to a subject to overcome tumor-specific tolerance in the subject. If desired, the tumor cell can be transfected to express a combination of peptides. For example, tumor cells obtained from a patient can be transfected ex vivo with an expression

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vector directing the expression of a peptide having B7-2-like activity alone, or in conjunction with a peptide having B7-1-like activity and/or B7-3-like activity. The transfected tumor cells are returned to the patient to result in expression of the peptides on the surface of the transfected cell. Alternatively, gene therapy techniques can be used to target a tumor cell for transfection in vivo.

The presence of the peptide of the present invention having the activity of a B lymphocyte antigen(s) on the surface 10 of the tumor cell provides the necessary costimulation signal to T cells to induce a T cell mediated immune response against the transfected tumor cells. In addition, tumor cells which lack MHC class I or MHC class II molecules, or which fail to reexpress sufficient amounts of MHC class I or MHC class II 15 molecules, can be transfected with nucleic acid encoding all or a portion of (e.g., a cytoplasmic-domain truncated portion) of an MHC class I  $\alpha$  chain protein and  $\beta_2$  microglobulin protein or an MHC class II $\alpha$  chain protein and an MHC class II $\beta$  chain protein to thereby express MHC class I or MHC class II proteins 20 on the cell surface. Expression of the appropriate class I or class II MHC in conjunction with a peptide having the activity of a B lymphocyte antigen (e.g., B7-1, B7-2, B7-3) induces a T cell mediated immune response against the transfected tumor cell. Optionally, a gene encoding an antisense construct which 25 blocks expression of an MHC class II associated protein, such as the invariant chain, can also be cotransfected with a DNA encoding a peptide having the activity of a B lymphocyte antigen to promote presentation of tumor associated antigens and induce tumor specific immunity. Thus, the induction of a

T cell mediated immune response in a human subject may be sufficient to overcome tumor-specific tolerance in the subject.

The activity of a protein of the invention may, among other means, be measured by the following methods:

5 Suitable assays for thymocyte or splenocyte cytotoxicity include, without limitation, those described in: Protocols in Immunology, Ed by J. E. Coligan, A.M. Kruisbeek, D.H. Margulies, E.M. Shevach, W Strober, Pub. Greene Publishing Associates and Wiley-Interscience (Chapter 3, In Vitro assays for Mouse Lymphocyte Function 3.1-3.19; Chapter 7, Immunologic studies in Humans); Herrmann et al., Proc. Natl. Acad. Sci. USA 78:2488-2492, 1981; Herrmann et al., J. Immunol. 128:1968-1974, 1982; Handa et al., J. Immunol. 135:1564-1572, 1985; Takai et al., J. Immunol. 137:3494-3500, 1986; Takai et al., J. Immunol. 15 140:508-512, 1988; Herrmann et al., Proc. Natl. Acad. Sci. USA 78:2488-2492, 1981; Herrmann et al., J. Immunol. 128:1968-1974, 1982; Handa et al., J. Immunol. 135:1564-1572, 1985; Takai et al., J. Immunol. 137:3494-3500, 1986; Bowmanet al., J. Virology 61:1992-1998; Takai et al., J. Immunol. 140:508-512, 20 Bertagnolli et al., Cellular Immunology 133:327-341, 1991; Brown et al., J. Immunol. 153:3079-3092, 1994.

Assays for T-cell-dependent immunoglobulin responses and isotype switching (which will identify, among others, proteins that modulate T-cell dependent antibody responses and that 25 affect Th1/Th2 profiles) include, without limitation, those described in: Maliszewski, J. Immunol. 144:3028-3033, 1990; and Assays for B cell function: In vitro antibody production, Mond, J.J. and Brunswick, M. In Current Protocols in Immunology. J.E.e.a. Coligan eds. Vol 1 pp. 3.8.1-3.8.16, John

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Wiley and Sons, Toronto. 1994.

Mixed lymphocyte reaction (MLR) assays (which will identify, among others, proteins that generate predominantly Thl and CTL responses) include, without limitation, those 5 described in: Current Protocols in Immunology, Ed by J. E. Coligan, A.M. Kruisbeek, D.H. Margulies, E.M. Shevach, W Strober, Pub. Greene Publishing Associates and Wiley-Interscience (Chapter 3, In Vitro assays for Mouse Lymphocyte Function 3.1-3.19; Chapter 7, Immunologic studies in 10 Humans); Takai et al., J. Immunol. 137:3494-3500, 1986; Takai et al., J. Immunol. 140:508-512, 1988; Bertagnolli et al., J. Immunol. 149:3778-3783, 1992.

Dendritic cell-dependent assays (which will identify, among others, proteins expressed by dendritic cells that activate naive T-cells) include, without limitation, those 15 described in: Guery et al., J. Immunol. 134:536-544, 1995; Inaba et al., Journal of Experimental Medicine 173:549-559, 1991; Macatonia et al., Journal of Immunology 154:5071-5079, 1995; Porgador et al., Journal of Experimental Medicine 20 182:255-260, 1995; Nair et al., Journal of Virology 67:4062-4069, 1993; Huang et al., Science 264:961-965, 1994; Macatonia et al., Journal of Experimental Medicine 169:1255-1264, 1989; Bhardwaj et al., Journal of Clinical Investigation 94:797-807, 1994; and Inaba et al., Journal of Experimental Medicine 172:631-640, 1990.

Assays for lymphocyte survival/apoptosis (which will identify, among others, proteins that prevent apoptosis after superantigen induction and proteins that regulate lymphocyte homeostasis) include, without limitation, those described in:

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Darzynkiewicz et al., Cytometry 13:795-808, 1992; Gorczyca et al., Leukemia 7:659-670, 1993; Gorczyca et al., Cancer Research 53:1945-1951, 1993; Itoh et al., Cell 66:233-243, 1991; Zacharchuk, Journal of Immunology 145:4037-4045, 1990; Zamai et al., Cytometry 14:891-897, 1993; Gorczyca et al., International Journal of Oncology 1:639-648, 1992.

Assays for proteins that influence early steps of T-cell commitment and development include, without limitation, those described in: Antica et al., Blood 84:111-117, 1994; Fine et al., Cellular Immunology 155:111-122, 1994; Galy et al., Blood 85:2770-2778, 1995; Toki et al., Proc. Nat. Acad Sci. USA 88:7548-7551, 1991.

## Hematopoiesis Regulating Activity

A protein of the present invention may be useful in 15 regulation of hematopoiesis and, consequently, in the treatment of myeloid or lymphoid cell deficiencies. Even marginal biological activity in support of colony forming cells or of factor-dependent cell lines indicates involvement in regulating hematopoiesis, e.g. in supporting the growth and proliferation 20 of erythroid progenitor cells alone or in combination with other cytokines, thereby indicating utility, for example, in treating various anemias or for use in conjunction with irradiation/chemotherapy to stimulate the production of erythroid precursors and/or erythroid cells; in supporting the 25 growth and proliferation of myeloid cells such as granulocytes and monocytes/macrophages (i.e., traditional CSF activity) useful, for example, in conjunction with chemotherapy to prevent or treat consequent myelo-suppression; in supporting the growth and proliferation of megakaryocytes and consequently

of platelets thereby allowing prevention or treatment of various platelet disorders such as thrombocytopenia, and generally for use in place of or complimentary to platelet transfusions; and/or in supporting the growth and proliferation 5 of hematopoietic stem cells which are capable of maturing to any and all of the above-mentioned hematopoietic cells and therefore find therapeutic utility in various stem cell disorders (such as those usually treated with transplantation, including, without limitation, aplastic anemia and paroxysmal 10 nocturnal hemoglobinuria), as well as in repopulating the stem cell compartment post irradiation/chemotherapy, either in-vivo (i.e., or ex-vivo in conjunction with bone marrow or progenitor transplantation with peripheral cell transplantation (homologous or heterologous)) as normal cells 15 or genetically manipulated for gene therapy.

The activity of a protein of the invention may, among other means, be measured by the following methods:

Suitable assays for proliferation and differentiation of various hematopoietic lines are cited above.

Assays for embryonic stem cell differentiation (which will identify, among others, proteins that influence embryonic differentiation hematopoiesis) include, without limitation, those described in: Johansson et al. Cellular Biology 15:141-151, 1995; Keller et al., Molecular and Cellular Biology 13:473-486, 1993; McClanahan et al., Blood 81:2903-2915, 1993.

Assays for stem cell survival and differentiation (which will identify, among others, proteins that regulate lympho-hematopoiesis) include, without limitation, those described in: Methylcellulose colony forming assays, Freshney,

M.G. In Culture of Hematopoietic Cells. R.I. Freshney, et al. eds. Vol pp. 265-268, Wiley-Liss, Inc., New York, NY. 1994; Hirayama et al., Proc. Natl. Acad. Sci. USA 89:5907-5911, 1992; Primitive hematopoietic colony forming cells with high 5 proliferative potential, McNiece, I.K. and Briddell, R.A. In Culture of Hematopoietic Cells. R.I. Freshney, et al. eds. Vol pp. 23-39, Wiley-Liss, Inc., New York, NY. 1994; Neben et al., Experimental Hematology 22:353-359, 1994; Cobblestone area assay, Ploemacher, R.E. forming cell In Culture Hematopoietic Cells. R.I. Freshney, et al. eds. Vol pp. 1-21, Wiley-Liss, Inc., New York, NY. 1994; Long term bone marrow cultures in the presence of stromal cells, Spooncer, E., Dexter, M. and Allen, T. In Culture of Hematopoietic Cells. R.I. Freshney, et al. eds. Vol pp. 163-179, Wiley-Liss, Inc., 15 New York, NY. 1994; Long term culture initiating cell assay, Sutherland, H.J. In Culture of Hematopoietic Cells. R.I. Freshney, et al. eds. Vol pp. 139-162, Wiley-Liss, Inc., New York, NY. 1994.

#### Tissue Growth Activity

- A protein of the present invention also may have utility in compositions used for bone, cartilage, tendon, ligament and/or nerve tissue growth or regeneration, as well as for wound healing and tissue repair and replacement, and in the treatment of burns, incisions and ulcers.
- A protein of the present invention, which induces cartilage and/or bone growth in circumstances where bone is not normally formed, has application in the healing of bone fractures and cartilage damage or defects in humans and other animals. Such a preparation employing a protein of the

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invention may have prophylactic use in closed as well as open fracture reduction and also in the improved fixation of artificial joints. De novo bone formation induced by an osteogenic agent contributes to the repair of congenital, trauma induced, or oncologic resection induced craniofacial defects, and also is useful in cosmetic plastic surgery.

A protein of this invention may also be used in the treatment of periodontal disease, and in other tooth repair processes. Such agents may provide an environment to attract bone-forming cells, stimulate growth of bone-forming cells or induce differentiation of progenitors of bone-forming cells. A protein of the invention may also be useful in the treatment of osteoporosis or osteoarthritis, such as through stimulation of bone and/or cartilage repair or by blocking inflammation or processes of tissue destruction (collagenase activity, osteoclast activity, etc.) mediated by inflammatory processes.

Another category of tissue regeneration activity that may be attributable to the protein of the present invention is tendon/ligament formation. A protein of the present invention, 20 which induces tendon/ligament-like tissue or other tissue formation in circumstances where such tissue is not normally formed, has application in the healing of tendon or ligament tears, deformities and other tendon or ligament defects in humans and other animals. Such a preparation employing a 25 tendon/ligament-like inducing protein tissue prophylactic use in preventing damage to tendon or ligament tissue, as well as use in the improved fixation of tendon or ligament to bone or other tissues, and in repairing defects to tendon or ligament tissue. De novo tendon/ligament-like tissue

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formation induced by a composition of the present invention contributes to the repair of congenital, trauma induced, or other tendon or ligament defects of other origin, and is also useful in cosmetic plastic surgery for attachment or repair of tendons or ligaments. The compositions of the present invention may provide an environment to attract tendon- or stimulate growth of tendonligament-forming cells, ligament-forming cells, induce differentiation of progenitors of tendon- or ligament-forming cells, or induce growth of 10 tendon/ligament cells or progenitors ex vivo for return in vivo to effect tissue repair. The compositions of the invention may also be useful in the treatment of tendinitis, carpal tunnel ligament defects. syndrome and other tendon or The compositions may also include an appropriate matrix and/or sequestering agent as a carrier as is well known in the art.

15

The protein of the present invention may also be useful for proliferation of neural cells and for regeneration of nerve and brain tissue, i.e. for the treatment of central and peripheral nervous system diseases and neuropathies, as well as 20 mechanical and traumatic disorders, which involve degeneration, death or trauma to neural cells or nerve tissue. More specifically, a protein may be used in the treatment of diseases of the peripheral nervous system, such as peripheral neuropathy localized nerve injuries, peripheral and neuropathies, and central nervous system diseases, such as Alzheimer's, Parkinson's disease, Huntington's amyotrophic lateral sclerosis, and Shy-Drager syndrome. Further conditions which may be treated in accordance with the present invention include mechanical and traumatic disorders,

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such as spinal cord disorders, head trauma and cerebrovascular diseases such as stroke. Peripheral neuropathies resulting from chemotherapy or other medical therapies may also be treatable using a protein of the invention.

Proteins of the invention may also be useful to promote better or faster closure of non-healing wounds, including without limitation pressure ulcers, ulcers associated with vascular insufficiency, surgical and traumatic wounds, and the like.

It is expected that a protein of the present invention may also exhibit activity for generation or regeneration of other tissues, such as organs (including, for example, pancreas, liver, intestine, kidney, skin, endothelium), muscle (smooth, skeletal or cardiac) and vascular (including vascular endothelium) tissue, or for promoting the growth of cells comprising such tissues. Part of the desired effects may be by inhibition or modulation of fibrotic scarring to allow normal tissue to regenerate. A protein of the invention may also exhibit angiogenic activity.

20 A protein of the present invention may also be useful for gut protection or regeneration and treatment of lung or liver fibrosis, reperfusion injury in various tissues, and conditions resulting from systemic cytokine damage.

A protein of the present invention may also be useful for promoting or inhibiting differentiation of tissues described above from precursor tissues or cells; or for inhibiting the growth of tissues described above.

The activity of a protein of the invention may, among other means, be measured by the following methods:

Assays for tissue generation activity include, without limitation, those described in: International WO95/16035 cartilage, tendon); Publication No. (bone, Patent Publication No. WO95/05846 International 5 neuronal); International Patent Publication No. W091/07491 (skin, endothelium).

Assays for wound healing activity include, without limitation, those described in: Winter, Epidermal Wound Healing, pps. 71-112 (Maibach, HI and Rovee, DT, eds.), Year 10 Book Medical Publishers, Inc., Chicago, as modified by Eaglstein and Mertz, J. Invest. Dermatol 71:382-84 (1978).

## Activin/Inhibin Activity

A protein of the present invention may also exhibit inhibin-related activities. Inhibins activinorcharacterized by their ability to inhibit the release of follicle stimulating hormone (FSH), while activins and are characterized by their ability to stimulate the release of follicle stimulating hormone (FSH). Thus, a protein of the present invention, alone or in heterodimers with a member of 20 the inhibin  $\alpha$  family, may be useful as a contraceptive based on the ability of inhibins to decrease fertility in female mammals and decrease spermatogenesis in male mammals. Administration of sufficient amounts of other inhibins can induce infertility in these mammals. Alternatively, the protein of the invention, as a homodimer or as a heterodimer with other protein subunits of the inhibin- $\beta$  group, may be useful as a fertility inducing therapeutic, based upon the ability of activin molecules in stimulating FSH release from cells of the anterior pituitary. See, for example, United States Patent 4,798,885. A protein of

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the invention may also be useful for advancement of the onset of fertility in sexually immature mammals, so as to increase the lifetime reproductive performance of domestic animals such as cows, sheep and pigs.

5 The activity of a protein of the invention may, among other means, be measured by the following methods:

Assays for activin/inhibin activity include, without limitation, those described in: Vale et al., Endocrinology 91:562-572, 1972; Ling et al., Nature 321:779-782, 1986; Vale et al., Nature 321:776-779, 1986; Mason et al., Nature 318:659-663, 1985; Forage et al., Proc. Natl. Acad. Sci. USA 83:3091-3095, 1986.

## Chemotactic/Chemokinetic Activity

A protein of the present invention may have chemotactic or chemokinetic activity (e.g., act as a chemokine) for mammalian cells, including, for example, monocytes, fibroblasts, neutrophils, T-cells, mast cells, eosinophils, epithelial and/or endothelial cells. Chemotactic and chemokinetic proteins can be used to mobilize or attract a desired cell population to a desired site of action. Chemotactic or chemokinetic proteins provide particular advantages in treatment of wounds and other trauma to tissues, as well as in treatment of localized infections. For example, attraction of lymphocytes, monocytes or neutrophils to tumors or sites of infection may result in improved immune responses against the tumor or infecting agent.

A protein or peptide has chemotactic activity for a particular cell population if it can stimulate, directly or indirectly, the directed orientation or movement of such cell

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population. Preferably, the protein or peptide has the ability to directly stimulate directed movement of cells. Whether a particular protein has chemotactic activity for a population of cells can be readily determined by employing such protein or peptide in any known assay for cell chemotaxis.

The activity of a protein of the invention may, among other means, be measured by the following methods:

Assays for chemotactic activity (which will identify proteins that induce or prevent chemotaxis)consist of assays

10 that measure the ability of a protein to induce the migration of cells across a membrane as well as the ability of a protein to induce the adhesion of one cell population to another cell population. Suitable assays for movement and adhesion include, without limitation, those described in: Current Protocols in

15 Immunology, Ed by J.E. Coligan, A.M. Kruisbeek, D.H. Margulies, E.M. Shevach, W.Strober, Pub. Greene Publishing Associates and Wiley-Interscience (Chapter 6.12, Measurement of alpha and beta Chemokines 6.12.1-6.12.28; Taub et al. J. Clin. Invest. 95:1370-1376, 1995; Lind et al. APMIS 103:140-146, 1995; Muller et al Eur. J. Immunol. 25: 1744-1748; Gruber et al. J. of Immunol. 152:5860-5867, 1994; Johnston et al. J. of Immunol. 153: 1762-1768, 1994.

## Hemostatic and Thrombolytic Activity

A protein of the invention may also exhibit hemostatic or thrombolytic activity. As a result, such a protein is expected to be useful in treatment of various coagulation disorders (includinghereditary disorders, such as hemophilias) or to enhance coagulation and other hemostatic events in treating wounds resulting from trauma, surgery or other causes. A

protein of the invention may also be useful for dissolving or inhibiting formation of thromboses and for treatment and prevention of conditions resulting therefrom (such as, for example, infarction of cardiac and central nervous system 5 vessels (e.g., stroke).

The activity of a protein of the invention may, among other means, be measured by the following methods:

Assay for hemostatic and thrombolytic activity include, without limitation, those described in: Linet et al., J. Clin. Pharmacol. 26:131-140, 1986; Burdick et al., Thrombosis Res. 45:413-419, 1987; Humphrey et al., Fibrinolysis 5:71-79 (1991); Schaub, Prostaglandins 35:467-474, 1988.

## Receptor/Liquid Activity

A protein of the present invention may also demonstrate 15 activity as receptors, receptor ligands or inhibitors or agonists of receptor/ligand interactions. Examples of such receptors and ligands include, without limitation, cytokine receptors and their ligands, receptor kinases and their ligands, receptor phosphatases and their ligands, receptors involved in cell-cell interactions and their ligands (including without limitation, cellular adhesion molecules (such as selectins, integrins and their liqunds) and receptor/liqund pairs involved in antigen presentation, antigen recognition and development of cellular and humoral immune responses). Receptors and ligands are also useful for screening of potential peptide or small molecule inhibitors of the relevant receptor/ligand interaction. A protein of the present invention (including, without limitation, fragments receptors and ligands) may themselves be useful as inhibitors

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of receptor/ligand interactions.

The activity of a protein of the invention may, among other means, be measured by the following methods:

Suitable assays for receptor-ligand activity include

5 without limitation those described in:Current Protocols in
Immunology, Ed by J.E. Coligan, A.M. Kruisbeek, D.H. Margulies,
E.M. Shevach, W.Strober, Pub. Greene Publishing Associates and
Wiley-Interscience (Chapter 7.28, Measurement of Cellular
Adhesion under static conditions 7.28.1-7.28.22), Takai et al.,

10 Proc. Natl. Acad. Sci. USA 84:6864-6868, 1987; Bierer et al.,
J. Exp. Med. 168:1145-1156, 1988; Rosenstein et al., J. Exp.
Med. 169:149-160 1989; Stoltenborg et al., J. Immunol.
Methods 175:59-68, 1994; Stitt et al., Cell 80:661-670, 1995.

### Anti-Inflammatory Activity

15 Proteins of the present invention may also exhibit anti-inflammatory activity. The anti-inflammatory activity may be achieved by providing a stimulus to cells involved in the inflammatory response, by inhibiting or promoting cell-cell interactions (such as, for example, cell adhesion), by 20 inhibiting or promoting chemotaxis of cells involved in the inflammatory process, inhibiting or promoting cell extravasation, or by stimulating or suppressing production of other factors which more directly inhibit or promote an inflammatory response. Proteins exhibiting such activities can 25 be used to treat inflammatory conditions including chronic or acute conditions), including without limitation inflammation associated with infection (such as septic shock, sepsis or systemic inflammatory response syndrome (SIRS)), ischemia-reperfusion injury, endotoxin lethality, arthritis,

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complement-mediated hyperacute rejection, nephritis, cytokine or chemokine-induced lung injury, inflammatory bowel disease, Crohn's disease or resulting from over production of ytokines such as TNF or IL-1. Proteins of the invention may also be useful to treat anaphylaxis and hypersensitivity to an antigenic substance or material.

## Tumor Inhibition Activity

In addition to the activities described above for immunological treatment or prevention of tumors, a protein of the invention may exhibit other anti-tumor activities. A protein may inhibit tumor growth directly or indirectly (such as, for example, via ADCC). A protein may exhibit its tumor inhibitory activity by acting on tumor tissue or tumor precursor tissue, by inhibiting formation of tissues necessary to support tumor growth (such as, for example, by inhibiting angiogenesis), by causing production of other factors, agents or cell types which inhibit tumor growth, or by suppressing, eliminating or inhibiting factors, agents or cell types which promote tumor growth

## 20 Other Activities

A protein of the invention may also exhibit one or more of the following additional activities or effects: inhibiting the growth, infection or function of, or killing, infectious agents, including, without limitation, bacteria, viruses, fungi and other parasites; effecting (suppressing or enhancing) bodily characteristics, including, without limitation, height, weight, hair color, eye color, skin, fat to lean ratio or other tissue pigmentation, or organ or body part size or shape (such as, for example, breast augmentation or diminution, change in

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bone form or shape); effecting biorhythms or caricadic cycles or rhythms; effecting the fertility of male or female subjects; effecting the metabolism, catabolism, anabolism, processing, utilization, storage or elimination of dietary fat, lipid, 5 protein, carbohydrate, vitamins, minerals, cofactors or other nutritional factors or component(s); effecting behavioral characteristics, including, without limitation, appetite, libido, stress, cognition (including cognitive disorders), depression (including depressive disorders) and violent 10 behaviors; providing analgesic effects or other pain reducing effects; promoting differentiation and growth of embryonic stem cells in lineages other than hematopoietic lineages; hormonal or endocrine activity; in the case of enzymes, correcting deficiencies of the enzyme and treating deficiency-related 15 diseases; treatment of hyperproliferative disorders (such as, for example, psoriasis); immunoglobulin-like activity (such as, for example, the ability to bind antigens or complement); and the ability to act as an antigen in a vaccine composition to raise an immune response against such protein or another 20 material or entity which is cross-reactive with such protein.

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PCT/JP98/02445

# Sequence Table

	(2)	INF	ORMA	TION	FOR	SEQ	ID	NO:	1:							
5		(	i) S	EQUE	NCE	CHAR	ACTE	RIST	ICS:							
				(A)	LEN	GTH:	382									
				(B)	TYP	E: A	mino	aci	d							•
				(D)	TOP	OLOG	Y: L	inea	r							
		(	ii)	SEQU	ENCE	KIN	D: P	rote	in							
10		(	iii)	HYP	othe	TICA	L: N	0								
															,	
		(	vi)	ORIG	INAL	sou	RCE:									
				(A)	ORG.	ANIS	M: <i>H</i>	omo	sapi	ens .						
				(B)	CEL	L KI	ND:	Live	r							
15				(D)	CLO	NE N	AME:	HP0	1263							
		(:	xi)	SEQU	ENCE	DES	CRIP	TION	: SE	Q ID	NO:	1:				
	Met	Gly	Leu	Leu	Leu	Pro	Leu	Ala	Leu	Cys	Ile	Leu	Val	Leu	Сув	Су
20	1				5					10					15	
	Gly	Ala	Met	Ser	Pro	Pro	Gln	Leu	Ala	Leu	Asn	Pro	Ser	Ala	Leu	Le
				20					25					30		
	Ser	Arg		Cys	Asn	Asp	Ser	_	Val	Leu	Ala	Val		Gly	Phe	Al
			35					40					45			
25	Leu		Asp	Ile	Asn	Lys	-	Arg	Lys	Asp	Gly	-	Val	Leu	Arg	Le
		50					55					60				
		Arg	Val	Asn	Asp			Glu	Tyr	Arg	_	Gly	Gly	Leu	Gly	
	65		•			70					75					8
	Leu	Phe	Tyr	Leu	Thr	Leu	Asp	Val	Leu	Glu	Thr	Asp	Cys	His		Le
30					85					90					95	
	Arg	Lys	Lys		Trp	Gln	Asp	Cys	_	Met	Arg	Ile	Phe		Glu	Se
				100					105					110		
•	Val	Tyr	-	Gln	Cys	Lys	Ala		Phe	Tyr	Met	Asn		Pro	Ser	Ar
			115					120					125			
35	Val	Leu	Tyr	Leu	Ala	Ala	Tyr	Asn	Cvs	Thr	Leu	Arg	Pro	Val	Ser	Ly

Lys Lys Ile Tyr Met Thr Cys Pro Asp Cys Pro Ser Ser Ile Pro Thr

	Asp	Ser	Ser	Asn	His	Gln	Val	Leu	Glu	Ala	Ala	Thr	Glu	Ser	Leu	Ala
					165					170	)				175	
	Lys	Tyr	Asn	Asn	G1u	Asn	Thr	Ser	Lys	Glr	Tyr	Ser	Leu	Phe	Lys	Va]
				180					185					190		
5	Thr	Arg	Ala	Ser	Ser	Gln	Trp	Val	Val	Gly	Pro	Ser	Tyr	Phe	Val	Gli
			195					200					205			
	Tyr	Leu	Ile	Lys	Glu	Ser	Pro	Cys	Thr	Lys	Ser	Gln	Ala	Ser	Ser	Cys
		210					215					220				
	Ser	Leu	Gln	Ser	Ser	Asp	Ser	Va1	Pro	Val	Gly	Leu	Cys	Lys	Gly	Ser
10	225					230					235					240
	Leu	Thr	Arg	Thr	His	Trp	Glu	Lys	Phe	Val	Ser	Val	Thr	Cys	Asp	Phe
					245		•			250					255	
	Phe	Glu	Ser	Gln	Ala	Pro	Ala	Thr	Gly	Ser	Glu	Asn	Ser	Ala	Val	Asn
				260					265					270		
15	Gln	Lys	Pro	Thr	Asn	Leu	Pro	Lys	Val	Glu	Glu	Ser	Gln	Gln	Lys	Asn
			275					280					285			
	Thr	Pro	Pro	Thr	Asp	Ser	Pro	Ser	Lys	Ala	Gly	Pro	Arg	Gly	Ser	Va1
		290					295					300				
	G1n	Tyr	Leu	Pro	Asp	Leu	Asp	Asp	Lys	Asn	Ser	G1n	Glu	Lys	Gly	Pro
20	305					310					315					320
	Gln	Glu	Ala	Phe	Pro	Val	His	Leu	Asp	Leu	Thr	Thr	Asn	Pro	Gln	Gly
					325					330					335	
	Glu	Thr	Leu	Asp	Ile	Ser	Phe	Leu	Phe	Leu	Glu	Pro	Met	Glu	Glu	Lys
				340					345					350		
25	Leu	Val	Val	Leu	Pro	Phe	Pro	Lys	Glu	Lys	Ala	Arg	Thr	Ala	Glu	Cys
			355					360					365			
	Pro		Pro	Ala	Gln	Asn	Ala	Ser	Pro	Leu	Val	Leu	Pro	Pro		
	•	370					375					380				
(11)																

- (2) INFORMATION FOR SEQ ID NO: 2:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 317
    - (B) TYPE: Amino acid
- 35 (D) TOPOLOGY: Linear
  - (ii) SEQUENCE KIND: Protein
  - (iii) HYPOTHETICAL: No

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(vi)	ORIGINAL	SOURCE:
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- (A) ORGANISM: Homo sapiens
- (B) CELL KIND: Liver
- (D) CLONE NAME: HP01299

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#### (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 2:

Met Trp Leu Tyr Leu Ala Ala Phe Val Gly Leu Tyr Tyr Leu Leu His 10 Trp Tyr Arg Glu Arg Gln Val Val Ser His Leu Gln Asp Lys Tyr Val 25 30 Phe Ile Thr Gly Cys Asp Ser Gly Phe Gly Asn Leu Leu Ala Arg Gln 40 Leu Asp Ala Arg Gly Leu Arg Val Leu Ala Ala Cys Leu Thr Glu Lys 15 50 60 Gly Ala Glu Gln Leu Arg Gly Gln Thr Ser Asp Arg Leu Glu Thr Val 70 75 Thr Leu Asp Val Thr Lys Met Glu Ser Ile Ala Ala Ala Thr Gln Trp 90 20 Val Lys Glu His Val Gly Asp Arg Gly Leu Trp Gly Leu Val Asn Asn 100 105 Ala Gly Ile Leu Thr Pro Ile Thr Leu Cys Glu Trp Leu Asn Thr Glu 120 Asp Ser Met Asn Met Leu Lys Val Asn Leu Ile Gly Val Ile Gln Val 25 130 135 140 Thr Leu Ser Met Leu Pro Leu Val Arg Arg Ala Arg Gly Arg Ile Val 150 155 Asn Val Ser Ser Ile Leu Gly Arg Val Ala Phe Phe Val Gly Gly Tyr 165 170 30 Cys Val Ser Lys Tyr Gly Val Glu Ala Phe Ser Asp Ile Leu Arg Arg 185 Glu Ile Gln His Phe Gly Val Lys Ile Ser Ile Val Glu Pro Gly Tyr 195 200 Phe Arg Thr Gly Met Thr Asn Met Thr Gln Ser Leu Glu Arg Met Lys 35 215 Gln Ser Trp Lys Glu Ala Pro Lys His Ile Lys Glu Thr Tyr Gly Gln 230 235 Gln Tyr Phe Asp Ala Leu Tyr Asn Ile Met Lys Glu Gly Leu Leu Asn

86 245 250 255 Cys Ser Thr Asn Leu Asn Leu Val Thr Asp Cys Met Glu His Ala Leu 260 265 270 Thr Ser Val His Pro Arg Thr Arg Tyr Ser Ala Gly Trp Asp Ala Lys 280 Phe Phe Phe Ile Pro Leu Ser Tyr Leu Pro Thr Ser Leu Ala Asp Tyr 290 295 300 Ile Leu Thr Arg Ser Trp Pro Lys Pro Ala Gln Ala Val 305 310 315 10 (2) INFORMATION FOR SEQ ID NO: 3: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 296 15 (B) TYPE: Amino acid (D) TOPOLOGY: Linear (ii) SEQUENCE KIND: Protein (iii) HYPOTHETICAL: No 20 (vi) ORIGINAL SOURCE: (A) ORGANISM: Homo sapiens (B) CELL KIND: Liver (D) CLONE NAME: HP01347 25 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 3:

Met Ser Asp Ser Lys Glu Pro Arg Val Gln Gln Leu Gly Leu Leu Gly 10 Cys Leu Gly His Gly Ala Leu Val Leu Gln Leu Leu Ser Phe Met Leu 30 20 25 30 Leu Ala Gly Val Leu Val Ala Ile Leu Val Gln Val Ser Lys Val Pro Ser Ser Leu Ser Gln Glu Gln Ser Glu Gln Asp Ala Ile Tyr Gln Asn 55 35 Leu Thr Gln Leu Lys Ala Ala Val Gly Glu Leu Ser Glu Lys Ser Lys 65 70 Leu Gln Glu Ile Tyr Gln Glu Leu Thr Gln Leu Lys Ala Ala Val Gly 85 90

	Glu	Leu	Pro	Glu	Lys	Ser	Lys	Leu	Gln	Glu	Ile	Tyr	Gln	Glu	Leu	Thr
				100					105					110		
	Arg	Leu	Lys	Ala	Ala	Va1	Gly	Glu	Leu	Pro	Glu	Lys	Ser	Lys	Leu	Gln
			115					120					125			
5	Glu	Ile	Tyr	Gln	Glu	Leu	Thr	Arg	Leu	Lys	Ala	Ala	Val	Gly	Glu	Leu
		130					135					140				
	Pro	Glu	Lys	Ser	Lys	Leu	Gln	Glu	Ile	Tyr	Gln	Glu	Leu	Thr	Arg	Leu
	145					150					155					160
	Lys	Ala	Ala	Val	Gly	Glu	Leu	Pro	Glu	Lys	Ser	Lys	Leu	Gln	Glu	Ile
10					165					170					175	
	Tyr	Gln	Glu	Leu	Thr	Glu	Leu	Lys	Ala	Ala	Val	Gly	Glu	Leu	Pro	Glu
				180					185					190	•	
	Lys	Ser	Lys	Leu	Gln	Glu	Ile	Tyr	Gln	Glu	Leu	Thr	Gln	Leu	Lys	Ala
			195					200					205			
15	Ala	Val	Gly	Glu	Leu	Pro	Asp	Gln	Ser	Lys	Gln	Gln	Gln	I1e	Tyr	Gln
		210					215					220				
	Glu	Leu	Thr	Asp	Leu	Lys	Thr	Ala	Phe	Glu	Arg	Leu	Cys	Arg	His	Cys
	225					230				•	235					240
	Pro	Lys	Asp	Trp	Thr	Phe	Phe	Gln	Gly	Asn	Cys	Tyr	Phe	Met	Ser	Asn
20					245					250					255	
	Ser	Gln	Arg	Asn	Trp	His	Asp	Ser	Val	Thr	Ala	Cys	Gln	Glu	Val	Arg
				260					265					270		
	Ala	Gln	Leu	Val	Val	Ile	Lys	Thr-	Ala	Glu	Glu	Gln	Leu	Pro	Ala	Val
•			275					280					285			
25	Leu	Glu	Gln	Trp	Arg	Thr	Gln	Gln								
		290	•				295							•		

- (2) INFORMATION FOR SEQ ID NO: 4:
- 30 (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 197
  - (B) TYPE: Amino acid
  - (D) TOPOLOGY: Linear
  - (ii) SEQUENCE KIND: Protein
- 35 (iii) HYPOTHETICAL: No
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo sapiens

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1	(B)	CELL	KIND:	Stomach	cancer
	~ ~ /			O C OHIM CIT	CONCEL

(D) CLONE NAME: HP01440

#### (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 4:

5 Met Cys Thr Gly Lys Cys Ala Arg Cys Val Gly Leu Ser Leu Ile Thr 1 5 15 Leu Cys Leu Val Cys Ile Val Ala Asn Ala Leu Leu Val Pro Asn 20 25 Gly Glu Thr Ser Trp Thr Asn Thr Asn His Leu Ser Leu Gln Val Trp 10 35 40 45 Leu Met Gly Gly Phe Ile Gly Gly Leu Met Val Leu Cys Pro Gly Ile Ala Ala Val Arg Ala Gly Gly Lys Gly Cys Cys Gly Ala Gly Cys 15 75 Cys Gly Asn Arg Cys Arg Met Leu Arg Ser Val Phe Ser Ser Ala Phe 85 90 Gly Val Leu Gly Ala Ile Tyr Cys Leu Ser Val Ser Gly Ala Gly Leu 105 20 Arg Asn Gly Pro Arg Cys Leu Met Asn Gly Glu Trp Gly Tyr His Phe 115 120 125 Glu Asp Thr Ala Gly Ala Tyr Leu Leu Asn Arg Thr Leu Trp Asp Arg 135 Cys Glu Ala Pro Pro Arg Val Val Pro Trp Asn Val Thr Leu Phe Ser 25 145 150 155 Leu Leu Val Ala Ala Ser Cys Leu Glu Ile Val Leu Cys Gly Ile Gln 165 170 Leu Val Asn Ala Thr Ile Gly Val Phe Cys Gly Asp Cys Arg Lys Lys 180 185 190 30 Gln Asp Thr Pro His

- (2) INFORMATION FOR SEQ ID NO: 5:
  - (i) SEQUENCE CHARACTERISTICS:
- 35 (A) LENGTH: 221

- (B) TYPE: Amino acid
- (D) TOPOLOGY: Linear
- (ii) SEQUENCE KIND: Protein

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## (iii) HYPOTHETICAL: No

## (vi) ORIGINAL SOURCE:

- (A) ORGANISM: Homo sapiens
- 5 (B) CELL KIND: Stomach cancer
  - (D) CLONE NAME: HP01526

# (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 5:

. 10	Met	Glu	Ala	G1 y	Gly	Phe	Leu	Asp	Ser	Leu	Ile	Tyr	Gly	Ala	Сув	Val
	1				5					10					15	
	Val	Phe	Thr	Leu	Gly	Met	Phe	Ser	Ala	Gly	Leu	Ser	Asp	Leu	Arg	His
	٠			20					25					30		
	Met	Arg	Met	Thr	Arg	Ser	Val	Asp	Asn	Val	Gln	Phe	Leu	Pro	Phe	Leu
15			35					40					45			
	Thr	Thr	Glu	Val	Asn	Asn	Leu	Gly	Trp	Leu	Ser	Tyr	Gly	Ala	Leu	Lys
		50					55	•				60				
	Gly	Asp	Gly	Ile	Leu	Ile	Val	Va1	Asn	Thr	Va1	Gly	Ala	Ala	Leu	Gln
	65					70					75					80
20	Thr	Leu	Tyr	Ile	Leu	Ala	Tyr	Leu	His	Tyr	Cys	Pro	Arg	Lys	Arg	Val
					85					90					95	
	Val	Leu	Leu	Gln	Thr	Ala	Thr	Leu	Leu	G1y	Va1	Leu	Leu	Leu	Gly	Tyr
				100					105					110		
	Gly	Tyr	Phe	Trp	Leu	Leu	Val	Pro	Asn	Pro	Glu	Ála	Arg	Leu	Gln	Gln
25			115					120					125			
	Leu	Gly	Leu	Phe	Cys	Ser	Val	Phe	Thr	Ile	Ser	Met	Tyr	Leu	Ser	Pro
		130					135					140				
	Leu	Ala	Asp	Leu	Ala	Lys	Val	Ile	Gln	Thr	Lys	Ser	Thr	G1n	Cys	Leu
	145					150					155					160
30	Ser	Tyr	Pro	Leu	Thr	Ile	Ala	Thr	Leu	Leu	Thr	Ser	Ala	Ser	Trp	Cys
					165					170					175	
	Leu	Tyr	Gly	Phe	Arg	Leu	Arg	Asp	Pro	Tyr	Ile	Met	Val	Ser	Asn	Phe
				180					185					190		
	Pro	Gly	Ile	Val	Thr	Ser	Phe	Ile	Arg	Phe	Trp	Leu	Phe	Trp	Lys	Tyr
35			195					200					205			
	Pro	Gln	Glu	Gln	Asp	Arg	Asn	Tyr	Trp	Leu	Leu	Gln	Thr			
		210					215					220				

										90						
	(2)	INF	ORMA	TION	FOR	SEQ	ID	NO:	6:							
		(	i) S	EQUE	NCE	CHAR	ACTE	RIST	ics:							
				(A)	LEN	GTH:	251									
				(B)	TYP	E: A	mino	aci	d			•				
5				(D)	TOP	OLOG	Y: L	inea	r							
		(	ii)	SEQU	ENCE	KIN	D: P	rote	in							
		(	iii)	HYP	othe	TICA	L: N	0								
		,														
		(	vi)	ORIG	INAL	SOU	RCE:									
10				(A)	ORG	ANIS	M: <i>H</i>	ото	sapi	ens						
				(B)	CEL	L KI	ND:	Stom	ach	canc	er					
•				(D)	CLO	NE N	AME:	HP1	0230							
		(:	xi)	SEQU.	ENCE	DES	CRIP	TION	: SE	Q ID	NO:	6:				
15		_														
		Ser	Asp	Ile		Asp	Trp	Phe	Arg		Ile	Pro	Ala	Ile		Arg
	_ 1	_			5					10				_	15	
	Tyr	Trp	Phe	Ala	Ala	Thr	Val	Ala		Pro	Leu	Val	Gly	-	Leu	Gly
. 20	T 044	T1.	C	20	41.		*	DL -	25	m	<b>D</b>	01	41-	30	T	m
20	Leu	ire	35	Pro	Ala	Tyr	Leu		Leu	Trp	Pro	GIu		Pne	Leu	Tyr
	Ara	Pho		Ile	ሞሥክ	4=0	Dro	40 T10	ጥሎ 🕶	۸1.	<b>ም</b> ኤ =	Dho	45	Dha	Pro	Vo 1
	ME	50	GIII	116	пр	urg	55	116	1111	VIG	1111	60	ıyı	riie	rio	VAI
	Glv		Glv	Thr	Glv	Phe		Tvr	I.eu	Val	Asn	-	Tvr	Phe	Leu	Tvr
25	65		,	•	,	70		-,-	200		75	200	-,-			80
	Gln	Tyr	Ser	Thr	Arg		Glu	Thr	G1v	Ala		Asp	Gly	Arg	Pro	Ala
		•			85					90			,	Ü	95	
	Asp	Tyr	Leu	Phe	Met	Leu	Leu	Phe	Asn	Trp	Ile	Cys	Ile	Val	Ile	Thr
				100					105					110		
30	Gly	Leu	Ala	Met	Asp	Met	Gln	Leu	Leu	Met	Ile	Pro	Leu	Ile	Met	Ser
			115					120					125			
	Val	Leu	Tyr	Val	Trp	Ala	Gln	Leu	Asn	Arg	Asp	Met	Ile	Val	Ser	Phe
		130					135					140				
	Trp	Phe	G1y	Thr	Arg	Phe	Lys	Ala	Cys	Tyr	Leu	Pro	Trp	Val	Ile	Leu

150

Gly Phe Asn Tyr Ile Ile Gly Gly Ser Val Ile Asn Glu Leu Ile Gly

Asn Leu Val Gly His Leu Tyr Phe Phe Leu Met Phe Arg Tyr Pro Met

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				180					185					190		
	Asp	Leu	Gly	Gly	Arg	Asn	Phe	Leu	Ser	Thr	Pro	G1n	Phe	Leu	Tyr	Arg
			195					200					205	1		
	Trp	Leu	Pro	Ser	Arg	Arg	Gly	Gly	Va1	Ser	Gly	Phe	G1y	Val	Pro	Pro
5		210					215					220				
	Ala	Ser	Met	Arg	Arg	Ala	Ala	Asp	Gln	Asn	Gly	Gly	Gly	Gly	Arg	His
	225					230					235					240
	Asn	Trp	Gly	Gln	Gly	Phe	Arg	Leu	Gly	Asp	Gln					
					245					250					•	
10						•										
	(2)	INF	ORMA!	rion	FOR	SEQ	ID 1	NO:	7:							
		(:	i) SI	EQUE	NCE (	CHAR	ACTE	RIST	ICS:							
				(A)	LEN	GTH:	106					٠				
15				(B)	TYP	E: Ar	nino	acio	i							
				(D)	TOP	OLOG	C: L:	inear	r							
		(3	ii) S	EQU	ENCE	KIN	): P1	rote:	in							
		( :	lii)	HYP	THE:	ricai	L: No	)								
20		(1	7i) (													
					ORGA	•			-							
					CELI			_	ermo:	id c	arcin	noma				
					CELI											
				(D)	CLO	VE NA	ME:	HP10	389							
25																
		( )	ci) S	EQUI	ENCE	DESC	CRIPT	CION:	: SEC	) ID	NO:	7:				
	<b>W</b> -4	41.	M1	<b>5</b>	01	5	**. *	~-	_			_	_,		<b>5</b>	•
		ALA	rnr	Pro		Pro	Val	TTE	Pro		Val	Pro	Phe	Glu		Ser
20	1	D	D	17 - 1	5	01	01	• -	•	10	<b></b>		_		15	<b>~</b>
30	Lys	Pro	Pro		TIE	GIU	GIÀ	Leu		Pro	Thr	Val	Tyr	Arg	Asn	Pro
	C1	C	Db -	20	01	•	<b>D</b> L -	77 - <b>1</b>	25		<b>m</b> 1		0.1	30	D	.,
	GIU	ser		ras	GIU	Lys	Pne		Arg	Lys	Thr	Arg		Asn	Pro	VAI
	Vo 1	Des	35 T1-	01	0	T	41-	40	41.	. 1			45	<b>.</b>	01.	•
35	AGT		TTE	ота	СУS	Leu		inr	ATS	ATB	A18		rnr	Tyr	GIÀ	ьeu
<i>J J</i>	T***	50	Dha	n 1 ~	۸	C1	55	C =	C1-	A	C	60	1	Wa -	Mo+	A
	Tyr	oer	rne	115	Arg	GIÀ	ASN	ser	GID	Arg	ser	GIN	Leu	Met	met	Arg

Thr Arg Ile Ala Ala Gln Gly Phe Thr Val Ala Ala Ile Leu Leu Gly

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Leu Ala Val Thr Ala Met Lys Ser Arg Pro 100 105

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- (2) INFORMATION FOR SEQ ID NO: 8:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 78

. 10

- (B) TYPE: Amino acid
- (D) TOPOLOGY: Linear
- (ii) SEQUENCE KIND: Protein
- (iii) HYPOTHETICAL: No
- 15 (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo sapiens
  - (B) CELL KIND: Stomach cancer
  - (D) CLONE NAME: HP10408
- 20 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 8:

Met Gly Ser Gly Leu Pro Leu Val Leu Leu Leu Thr Leu Leu Gly Ser

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Ser His Gly Thr Gly Pro Gly Met Thr Leu Gln Leu Lys Leu Lys Glu

25

20

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Ser Phe Leu Thr Asn Ser Ser Tyr Glu Ser Ser Phe Leu Glu Leu Leu

35

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Glu Lys Leu Cys Leu Leu Leu His Leu Pro Ser Gly Thr Ser Val Thr
50 55 60

30 Leu His His Ala Arg Ser Gln His His Val Val Cys Asn Thr

65 70 75

- (2) INFORMATION FOR SEQ ID NO: 9:
- 35 (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 314
  - (B) TYPE: Amino acid
  - (D) TOPOLOGY: Linear

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(ii) SEQUENC	E KIND	Protein
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(iii) HYPOTHETICAL: No

#### (vi) ORIGINAL SOURCE:

5 (A) ORGANISM: Homo sapiens

- (B) CELL KIND: Stomach cancer
- (D) CLONE NAME: HP10412

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 9: 10 Met Val Ala Pro Val Trp Tyr Leu Val Ala Ala Ala Leu Leu Val Gly Phe Ile Leu Phe Leu Thr Arg Ser Arg Gly Arg Ala Ala Ser Ala Gly 25 15 Gln Glu Pro Leu His Asn Glu Glu Leu Ala Gly Ala Gly Arg Val Ala 35 40 45 Gln Pro Gly Pro Leu Glu Pro Glu Glu Pro Arg Ala Gly Gly Arg Pro Arg Arg Arg Arg Asp Leu Gly Ser Arg Leu Gln Ala Gln Arg Arg Ala 20 70 75 Gln Arg Val Ala Trp Ala Glu Ala Asp Glu Asn Glu Glu Glu Ala Val Ile Leu Ala Gln Glu Glu Glu Gly Val Glu Lys Pro Ala Glu Thr His 105 25 Leu Ser Gly Lys Ile Gly Ala Lys Lys Leu Arg Lys Leu Glu Glu Lys 115 120 125 Gln Ala Arg Lys Ala Gln Arg Glu Ala Glu Glu Ala Glu Arg Glu Glu 135 Arg Lys Arg Leu Glu Ser Gln Arg Glu Ala Glu Trp Lys Lys Glu Glu 30 145 150 Glu Arg Leu Arg Leu Glu Glu Glu Glu Glu Glu Glu Glu Arg Lys 170 Ala Arg Glu Glu Gln Ala Gln Arg Glu His Glu Glu Tyr Leu Lys Leu 180 185 35 Lys Glu Ala Phe Val Val Glu Glu Glu Gly Val Gly Glu Thr Met Thr 200 Glu Glu Gln Ser Gln Ser Phe Leu Thr Glu Phe Ile Asn Tyr Ile Lys 215 220

94

Gln Ser Lys Val Val Leu Leu Glu Asp Leu Ala Ser Gln Val Gly Leu 240 235 225 230 Arg Thr Gln Asp Thr Ile Asn Arg Ile Gln Asp Leu Leu Ala Glu Gly 245 250 Thr Ile Thr Gly Val Ile Asp Asp Arg Gly Lys Phe Ile Tyr Ile Thr 260 265 Pro Glu Glu Leu Ala Ala Val Ala Asn Phe Ile Arg Gln Arg Gly Arg 275 280 285 Val Ser Ile Ala Glu Leu Ala Gln Ala Ser Asn Ser Leu Ile Ala Trp 10 295 300 Gly Arg Glu Ser Pro Ala Gln Ala Pro Ala 305 310

- (2) INFORMATION FOR SEQ ID NO: 10: 15
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 195
    - (B) TYPE: Amino acid
    - (D) TOPOLOGY: Linear
- 20 (ii) SEQUENCE KIND: Protein
  - (iii) HYPOTHETICAL: No
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo sapiens
- 25 (B) CELL KIND: Stomach cancer
  - (D) CLONE NAME: HP10413
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 10:
- 30 Met Ala Ala Glu Asp Val Val Ala Thr Gly Ala Asp Pro Ser Asp Leu 10 Glu Ser Gly Gly Leu Leu His Glu Ile Phe Thr Ser Pro Leu Asn Leu 25 Leu Leu Cly Leu Cys Ile Phe Leu Leu Tyr Lys Ile Val Arg Gly 35 35 40 Asp Gln Pro Ala Ala Ser Gly Asp Ser Asp Asp Glu Pro Pro Pro 55

Leu Pro Arg Leu Lys Arg Arg Asp Phe Thr Pro Ala Glu Leu Arg Arg

95

70 65 75 80 Phe Asp Gly Val Gln Asp Pro Arg Ile Leu Met Ala Ile Asn Gly Lys 85 90 Val Phe Asp Val Thr Lys Gly Arg Lys Phe Tyr Gly Pro Glu Gly Pro 5 105 Tyr Gly Val Phe Ala Gly Arg Asp Ala Ser Arg Gly Leu Ala Thr Phe 115 Cys Leu Asp Lys Glu Ala Leu Lys Asp Glu Tyr Asp Asp Leu Ser Asp 135 140 10 Leu Thr Ala Ala Gln Gln Glu Thr Leu Ser Asp Trp Glu Ser Gln Phe 145 150 160 155 Thr Phe Lys Tyr His His Val Gly Lys Leu Leu Lys Glu Gly Glu Glu 165 170 Pro Thr Val Tyr Ser Asp Glu Glu Glu Pro Lys Asp Glu Ser Ala Arg 15 185 190 Lys Asn Asp 195

- 20 (2) INFORMATION FOR SEQ ID NO: 11:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 462
    - (B) TYPE: Amino acid
    - (D) TOPOLOGY: Linear
- 25 (ii) SEQUENCE KIND: Protein
  - (iii) HYPOTHETICAL: No
    - (vi) ORIGINAL SOURCE:
      - (A) ORGANISM: Homo sapiens
- 30 (B) CELL KIND: Stomach cancer
  - (D) CLONE NAME: HP10415
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 11:
- 35 Met Leu Asp Phe Ala Ile Phe Ala Val Thr Phe Leu Leu Ala Leu Val

  1 5 10 15

  Gly Ala Val Leu Tyr Leu Tyr Pro Ala Ser Arg Gln Ala Ala Gly Ile
  20 25 30

	Pro	Gly	Ile	Thr	Pro	Thr	Glu	Glu	Lys	Asp	Gly	Asn	Leu	Pro	Asp	Ile
			35					40					45			
	Val	Asn	Ser	Gly	Ser	Leu	His	Glu	Phe	Leu	Val	Asn	Leu	His	Glu	Arg
		50					55					60				
5	Tyr	Gly	Pro	Val	Val	Ser	Phe	Trp	Phe	Gly	Arg	Arg	Leu	Val	Val	Ser
	65					70					75					80
	Leu	Gly	Thr	Val	Asp	Val	Leu	Lys	Gln	His	Ile	Asn	Pro	Asn	Lys	Thr
					85					90					95	
	Leu	Asp	Pro	Phe	Glu	Thr	Met	Leu	Lys	Ser	Leu	Leu	Arg	Tyr	Gln	Ser
10				100					105					110		
	Gly	Gly	Gly	Ser	Val	Ser	Glu	Asn	His	Met	Arg	Lys	Lys	Leu	Tyr	Glu
			115					120					125			
	Asn	Gly	Val	Thr	Asp	Ser	Leu	Lys	Ser	Asn	Phe	Ala	Leu	Leu	Leu	Lys
		130					135					140				
L 5	Leu	Ser	Glu	Glu	Leu	Leu	Asp	Lys	Trp	Leu	Ser	Tyr	Pro	Glu	Thr	Gln
	145					150					155					160
	His	Val	Pro	Leu	Ser	Gln	His	Met	Leu	Gly	Phe	Ala	Met	Lys	Ser	Val
					165					170					175	
	Thr	Gln	Met	Val	Met	Gly	Ser	Thr	Phe	Glu	Asp	Asp	Gln	Glu	Val	Ile
20				180					185					190		
	Arg	Phe	Gln	Lys	Asn	His	Gly	Thr	Val	Trp	Ser	Glu	Ile	Gly	Lys	Gly
			195					200					205			
	Phe	Leu	Asp	Gly	Ser	Leu	Asp	Lys	Asn	Met	Thr	Arg	Lys	Lys	Gln	Tyr
		210					215					220				
25	Glu	Asp	Ala	Leu	Met	Gln	Leu	Glu	Ser	Val	Leu	Arg	Asn	Ile	Ile	Lys
	225					230					235					240
	Glu	Arg	Lys	Gly	Arg	Asn	Phe	Ser	Gln	His	Ile	Phe	Ile	Asp	Ser	Leu
					245					250					255	
	Val	Gln	Gly	Asn	Leu	Asn	Asp	Gln	Gln	Ile	Leu	Glu	Asp	Ser	Met	Ile
30				260					265					270		
	Phe	Ser	Leu	Ala	Ser	Cys	Ile	Ile	Thr	Ala	Lys	Leu	Cys	Thr	Trp	Ala
			275					280					285			
	Ile	Cys	Phe	Leu	Thr	Thr	Ser	Glu	Glu	Val	Gln	Lys	Lys	Leu	Tyr	G1u
		290					295					300				
35	Glu	lle	Asn	Gln	Val	Phe	Gly	Asn	Gly	Pro	Val	Thr	Pro	Glu	Lys	Ile
	305					310					315					320
	Glu	Gln	Leu	Arg	Tyr	Cys	Gln	His	Val	Leu	Cys	G1u	Thr	Val	Arg	Thr
					325					330					335	

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							•									
	Ala	Lys	Leu	Thr	Pro	Val	Ser	Ala	Gln	Leu	Gln	Asp	Ile	Glu	Gly	Ly
				340					345					350		
	Ile	Asp	Arg	Phe	Ile	Ile	Pro	Arg	Glu	Thr	Leu	Val	Leu	Tyr	Ala	Le
			355					360					365			
5	Gly	Val	Val	Leu	Gln	Asp	Pro	Asn	Thr	Trp	Pro	Ser	Pro	His	Lys	Ph
		370					375					380				
	Asp	Pro	Asp	Arg	Phe	Asp	Asp	Glu	Leu	Val	Met	Lys	Thr	Phe	Ser	Se
	385					390					395					40
	Leu	Gly	Phe	Ser	-	Thr	Gln	Glu	Cys	Pro	Glu	Leu	Arg	Phe	Ala	Ty
10					405					410			•		415	
	Met	Val	Thr		Val	Leu	Leu	Ser		Leu	Val	Lys	Arg		His	Le
				420				_	425					430		
	Leu	Ser		Glu	Gly	Gln	Val		Glu	Thr	Lys	Tyr		Leu	Val	Th
	_	_	435				_	440			_		445	_		
15	Ser	Ser	Arg	Glu	Glu	Ala	-	lle	Thr	Val	Ser		Arg	Tyr		
		450					455					460				
	(2)	INFO	NDMA'	יד חא	FAD	SEO.	TD N	in. 1	٠.							
20	(2)		.) SE			-										
		``	.,	-		STH:										
						E: An		ació	l			•				
						LOGY										
		( i	.i) S	EQUE	ENCE	KINI	): P1	otei	n							
25		i)	.ii)	HYPO	THE	CICAL	.: No	)								
		( v	ri) (	RIGI	NAL	SOUF	RCE:	•								
				(A)	ORGA	ANISM	1: H	omo s	sapie	ens						
				(B)	CELI	L KIN	ID: S	toma	ch c	ance	r					
30				(D)	CLO	NE NA	ME:	HP10	419							
		( x	i) S	EQUE	ENCE	DESC	RIPI	:NOI	SEQ	ID	NO:	12:				
	Met	Gly	Ala	Ala	Val	Phe	Phe	Gly	Cys	Thr	Phe	Val	Ala	Phe	Gly	Pro
35	1				5					10					15	
	Ala	Phe	Ala	Leu	Phe	Leu	Ile	Thr	Val	Ala	Gly	Asp	Pro	Leu	Arg	Va]
				20					25					30		

Ile Ile Leu Val Ala Gly Ala Phe Phe Trp Leu Val Ser Leu Leu Leu

			35					40					45			
	Ala	Ser	Val	Val	Trp	Phe	Ile	Leu	Val	His	Val	Thr	Asp	Arg	Ser	Asj
		50					55					60				
	Ala	Arg	Leu	Gln	Tyr	Gly	Leu	Leu	Ile	Phe	Gly	Ala	Ala	Val	Ser	Va]
5	65					70					75					80
	Leu	Leu	Gln	Glu	Val	Phe	Arg	Phe	Ala	Tyr	Tyr	Lys	Leu	Leu	Lys	Lys
					85					90					95	
	Ala	Asp	Glu	G1y	Leu	Ala	Ser	Leu	Ser	Glu	Asp	Gly	Arg	Ser	Pro	Ilε
				100	•				105				•	110		
10	Ser	Ile	Arg	Gln	Met	Ala	Tyr	Val	Ser	Gly	Leu	Ser	Phe	Gly	Ile	Ile
			115					120					125			
	Ser	Gly	Val	Phe	Ser	Val	Ile	Asn	Ile	Leu	Ala	Asp	Ala	Leu	Gly	Pro
		130					135					140				
	Gly	Val	Val	Gly	Ile	His	G1y	Asp	Ser	Pro	Tyr	Tyr	Phe	Leu	Thr	Ser
15	145					150					155					160
	Ala	Phe	Leu	Thr		Ala	Ile	Ile	Leu	Leu	His	Thr	Phe	Trp	Gly	Val
					165					170					175	
	Val	Phe	Phe		Ala	Cys	Glu	Arg		Arg	Tyr	Trp	Ala	Leu	Gly	Leu
•				180		_			185					190		
20	Val	Val		Ser	His	Leu	Leu		Ser	Gly	Leu	Thr		Leu	Asn	Pro
	<b></b>		195			_		200		_			205			
			GIU	Ala	ser	Leu		Pro	Ile	Tyr	Ala		Thr	Val	Ser	Met
		210	<b>m</b>	41.	Di-	<b>~1</b> -	215		0.1	-1	_	220		_		
25		Leu	Trp	ALA	rne		Tnr	ALA	GIÀ	Gly		Leu	Arg	Ser	IIe	
25	225	C 0 ==	Tau	Tau	0	230	A				235					240
	Arg	ser	ren	Leu	Cys	гàг	Asp									
					245											

- 30 (2) INFORMATION FOR SEQ ID NO: 13:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 113
    - (B) TYPE: Amino acid
    - (D) TOPOLOGY: Linear
- 35 (ii) SEQUENCE KIND: Protein
  - (iii) HYPOTHETICAL: No
  - (vi) ORIGINAL SOURCE:

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(A) ORGANISM: Homo sapiens

(B) CELL KIND: Stomach cancer

(D) CLONE NAME: HP10424

5 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 13:

Met Asn Phe Tyr Leu Leu Leu Ala Ser Ser Ile Leu Cys Ala Leu Ile

5 10 1

Val Phe Trp Lys Tyr Arg Arg Phe Gln Arg Asn Thr Gly Glu Met Ser

10 20 25 30

Ser Asn Ser Thr Ala Leu Ala Leu Val Arg Pro Ser Ser Gly Leu

35 40 45

Ile Asn Ser Asn Thr Asp Asn Asn Leu Ala Val Tyr Asp Leu Ser Arg

50 55 60

15 Asp Ile Leu Asn Asn Phe Pro His Ser Ile Ala Arg Gln Lys Arg Ile

65 70 75 80

Leu Val Asn Leu Ser Met Val Glu Asn Lys Leu Val Glu Leu Glu His
85 90 95

Thr Leu Leu Ser Lys Gly Phe Arg Gly Ala Ser Pro His Arg Lys Ser

20 100 105 110

Thr

- (2) INFORMATION FOR SEQ ID NO: 14:
- 25 (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 365
  - (B) TYPE: Amino acid
  - (D) TOPOLOGY: Linear
  - (ii) SEQUENCE KIND: Protein
- 30 (iii) HYPOTHETICAL: No
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo sapiens
    - (B) CELL KIND: Epidermoid carcinoma
- 35 (C) CELL LINE: KB
  - (D) CLONE NAME: HP10428
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 14:

	Met	Gly	Arg	Trp	Ala	Leu	Asp	Val	Ala	Phe	Leu	Trp	Lys	Ala	Val	Leu
	1				5					10					15	
	Thr	Leu	Gly	Leu	Val	Leu	Leu	Tyr	Tyr	Cys	Phe	Ser	Ile	Gly	Ile	Thr
				20					25					30		
5	Phe	Tyr	Asn	Lys	Trp	Leu	Thr	Lys	Ser	Phe	His	Phe	Pro	Leu	Phe	Met
			35					40					45			
	Thr	Met	Leu	His	Leu	Ala	Va1	Ile	Phe	Leu	Phe	Ser	Ala	Leu	Ser	Arg
		50					55					60				
	Ala	Leu	Val	Gln	Cys	Ser	Ser	His	Arg	Ala	Arg	Val	Val	Leu	Ser	Trp
10	65					70					75					80
	Ala	Asp	Tyr	Leu	Arg	Arg	Val	Ala	Pro	Thr	Ala	Leu	Ala	Thr	Ala	Leu
					85					90					95	
	Asp	Val	Gly	Leu	Ser	Asn	Trp	Ser	Phe	Leu	Tyr	Val	Thr	Val	Ser	Leu
				100					105					110		
15	Tyr	Thr	Met	Thr	Lys	Ser	Ser	Ala	Val	Leu	Phe	Ile	Leu	Ile	Phe	Ser
			115					120					125			
	Leu	Ile	Phe	Lys	Leu	Glu	Glu	Leu	Arg	Ala	Ala	Leu	Val	Leu	Val	Val
		130					135					140				
	Leu	Leu	Ile	Ala	Gly	Gly	Leu	Phe	Met	Phe	Thr	Tyr	Lys	Ser	Thr	Gln
20	145					150					155					160
	Phe	Asn	Val	Glu	Gly	Phe	Ala	Leu	Val	Leu	Gly	Ala	Ser	Phe	Ile	Gly
					165					170					175	
	Gly	Ile	Arg	Trp	Thr	Leu	Thr	Gln	Met	Leu	Leu	Gln	Lys	Ala	Glu	Leu
				180					185					190		
25	Gly	Leu	Gln	Asn	Pro	Ile	Asp	Thr	Met	Phe	His	Leu	Gln	Pro	Leu	Met
			195					200					205			
	Phe	Leu	Gly	Leu	Phe	Pro	Leu	Phe	Ala	Val	Phe	Glu	Gly	Leu	His	Leu
		210					215					220				
	Ser	Thr	Ser	Glu	Lys	Ile	Phe	Arg	Phe	Gln	Asp	Thr	Gly	Leu	Leu	Leu
30	225					230					235					240
	Arg	Val	Leu	Gly	Ser	Leu	Phe	Leu	Gly	Gly	Ile	Leu	Ala	Phe	-	Leu
					245					250					255	
	G1y	Phe	Ser	Glu	Phe	Leu	Leu	Val		Arg	Thr	Ser	Ser	Leu	Thr	Leu
				260					265					270		
35	Ser	Ile		Gly	Ile	Phe	Lys		Val	Cys	Thr	Leu	Leu	Leu	Ala	Ala
			275					280					285			
	His		Leu	Gly	Asp	Gln		Ser	Leu	Leu	Asn	-	Leu	Gly	Phe	Ala
		200					205					200				

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Leu Cys Leu Ser Gly Ile Ser Leu His Val Ala Leu Lys Ala Leu His 305 310 315 Ser Arg Gly Asp Gly Gly Pro Lys Ala Leu Lys Gly Leu Gly Ser Ser 325 330 Pro Asp Leu Glu Leu Leu Arg Ser Ser Gln Arg Glu Glu Gly Asp 345 Asn Glu Glu Glu Tyr Phe Val Ala Gln Gly Gln Gln 355 360 365 10 (2) INFORMATION FOR SEQ ID NO: 15: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 226 (B) TYPE: Amino acid 15 (D) TOPOLOGY: Linear (ii) SEQUENCE KIND: Protein (iii) HYPOTHETICAL: No (vi) ORIGINAL SOURCE: 20 (A) ORGANISM: Homo sapiens (B) CELL KIND: Stomach cancer (D) CLONE NAME: HP10429 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 15: 25 Met Pro Thr Thr Lys Lys Thr Leu Met Phe Leu Ser Ser Phe Phe Thr Ser Leu Gly Ser Phe Ile Val Ile Cys Ser Ile Leu Gly Thr Gln Ala 25 Trp Ile Thr Ser Thr Ile Ala Val Arg Asp Ser Ala Ser Asn Gly Ser 35 45 Ile Phe Ile Thr Tyr Gly Leu Phe Arg Gly Glu Ser Ser Glu Glu Leu 55 60 Ser His Gly Leu Ala Glu Pro Lys Lys Phe Ala Val Leu Glu Ile 35 65 70 75 Leu Asn Asn Ser Ser Gln Lys Thr Leu His Ser Val Thr Ile Leu Phe 90

Leu Val Leu Ser Leu Ile Thr Ser Leu Leu Ser Ser Gly Phe Thr Phe

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100 105 110 Tyr Asn Ser Ile Ser Asn Pro Tyr Gln Thr Phe Leu Gly Pro Thr Gly 115 120 125 Val Tyr Thr Trp Asn Gly Leu Gly Ala Ser Phe Val Phe Val Thr Met 135 140 Ile Leu Phe Val Ala Asn Thr Gln Ser Asn Gln Leu Ser Glu Glu Leu 150 155 Phe Gln Met Leu Tyr Pro Ala Thr Thr Ser Lys Gly Thr Thr His Ser 165 170 10 Tyr Gly Tyr Ser Phe Trp Leu Ile Leu Leu Val Ile Leu Leu Asn Ile 180 185 Val Thr Val Thr Ile Ile Ile Phe Tyr Gln Lys Ala Arg Tyr Gln Arg 195 200 205 Lys Gln Glu Gln Arg Lys Pro Met Glu Tyr Ala Pro Arg Asp Gly Ile 15 215 220 Leu Phe 225

- 20 (2) INFORMATION FOR SEQ ID NO: 16:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 129
    - (B) TYPE: Amino acid
    - (D) TOPOLOGY: Linear
- 25 (ii) SEQUENCE KIND: Protein
  - (iii) HYPOTHETICAL: No
    - (vi) ORIGINAL SOURCE:
      - (A) ORGANISM: Homo sapiens
- 30 (B) CELL KIND: Liver
  - (D) CLONE NAME: HP10432
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 16:
- 35 Met Ala Arg Gly Ser Leu Arg Arg Leu Leu Arg Leu Leu Val Leu Gly

  1 5 10 15

  Leu Trp Leu Ala Leu Leu Arg Ser Val Ala Gly Glu Gln Ala Pro Gly

  20 25 30

WO 98/55508

103

Thr Ala Pro Cys Ser Arg Gly Ser Ser Trp Ser Ala Asp Leu Asp Lys 35 40 Cys Met Asp Cys Ala Ser Cys Arg Ala Arg Pro His Ser Asp Phe Cys Leu Gly Cys Ala Ala Ala Pro Pro Ala Pro Phe Arg Leu Leu Trp Pro 70 75 Ile Leu Gly Gly Ala Leu Ser Leu Thr Phe Val Leu Gly Leu Leu Ser 85 Gly Phe Leu Val Trp Arg Arg Cys Arg Arg Glu Lys Phe Thr Thr 10 105 Pro Ile Glu Glu Thr Gly Gly Glu Gly Cys Pro Ala Val Ala Leu Ile 115 120 125 Gln 15 (2) INFORMATION FOR SEQ ID NO: 17: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 163 20 (B) TYPE: Amino acid (D) TOPOLOGY: Linear (ii) SEQUENCE KIND: Protein (iii) HYPOTHETICAL: No 25 (vi) ORIGINAL SOURCE: (A) ORGANISM: Homo sapiens (B) CELL KIND: Liver (D) CLONE NAME: HP10433 30 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 17: Met Arg Arg Leu Leu Ile Pro Leu Ala Leu Trp Leu Gly Ala Val Gly Val Gly Val Ala Glu Leu Thr Glu Ala Gln Arg Arg Gly Leu Gln Val 35 20 25 30 Ala Leu Glu Glu Phe His Lys His Pro Pro Val Gln Trp Ala Phe Gln 40

Glu Thr Ser Val Glu Ser Ala Val Asp Thr Pro Phe Pro Ala Gly Ile

104

50 60 55 Phe Val Arg Leu Glu Phe Lys Leu Gln Gln Thr Ser Cys Arg Lys Arg 65 70 75 Asp Trp Lys Lys Pro Glu Cys Lys Val Arg Pro Asn Gly Arg Lys Arg Lys Cys Leu Ala Cys Ile Lys Leu Gly Ser Glu Asp Lys Val Leu Gly 100 105 110 Arg Leu Val His Cys Pro Ile Glu Thr Gln Val Leu Arg Glu Ala Glu 120 10 Glu His Gln Glu Thr Gln Cys Leu Arg Val Gln Arg Ala Gly Glu Asp 135 Pro His Ser Phe Tyr Phe Pro Gly Gln Phe Ala Phe Ser Lys Ala Leu 145 150 155 Pro Arg Ser 15 (2) INFORMATION FOR SEQ ID NO: 18: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 193 20 (B) TYPE: Amino acid (D) TOPOLOGY: Linear (ii) SEQUENCE KIND: Protein (iii) HYPOTHETICAL: No 25 (vi) ORIGINAL SOURCE: (A) ORGANISM: Homo sapiens (B) CELL KIND: Stomach cancer (D) CLONE NAME: HP10480 30 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 18: Met Ile Arg Cys Gly Leu Ala Cys Glu Arg Cys Arg Trp Ile Leu Pro 1 5 10 Leu Leu Leu Ser Ala Ile Ala Phe Asp Ile Ile Ala Leu Ala Gly 35 Arg Gly Trp Leu Gln Ser Ser Asp His Gly Gln Thr Ser Ser Leu Trp 40

Trp Lys Cys Ser Gln Glu Gly Gly Gly Ser Gly Ser Tyr Glu Glu Gly

		50					55					60					
	Cys	Gln	Ser	Leu	Met	Glu	Tyr	Ala	Trp	Gly	Arg	Ala	Ala	Ala	Ala	Met	
	65					70					75					80	
	Leu	Phe	Cys	Gly	Phe	Ile	Ile	Leu	Val	Ile	Cys	Phe	Ile	Leu	Ser	Phe	
5					85					90					95		
	Phe	Ala	Leu	Cys	Gly	Pro	Gln	Met	Leu	Val	Phe	Leu	Arg	Val	Ile	Gly	
				100					105					110			
	Gly	Leu	Leu	Ala	Leu	Ala	Ala	Val	Phe	Gln	Ile	Ile	Ser	Leu	Val	Ile	
			115					120					125				
10	Tyr		Val	Lys	Tyr	Thr	Gln	Thr	Phe	Thr	Leu	His	Ala	Asn	Arg	Ala	
		130					135					140					
		Thr	Tyr	Ile	Tyr		Trp	Ala	Tyr	Gly		Gly	Trp	Ala	Ala		
	145					150					155					160	
	Ile	Ile	Leu	Ile	_	Cys	Ala	Phe	Phe		Cys	Cys	Leu	Pro	Asn	Tyr	
15	01	4	4	7	165	01	<b>.</b>	41	T	170	4	m	Dh.a	m	175	C	
	GIU	Asp	Asp		ren	GIÀ	Asn	AIA	•	Pro	Arg	Tyr	Pne	-	Thr	ser	
	Ala			180					185					190			
20																	
	(2)	INFO	RMAT	NOI	FOR	SEO	ID N	io: 1	9:								
	, ,					-		RISTI									
						TH:										,	
				(B)	TYPI	E: Nu	clei	c ac	id								
25				(C)	STRA	NDEI	NESS	: Do	uble	:			•				
				(D)	TOPO	LOGY	: Li	near	•								
		(i	i) S	EQUE	ENCE	KIND	: cI	NA t	o mR	NA							
		( v	i) (	RIGI	NAL	SOUR	CE:										
30				(A)	ORGA	NISM	l: Ho	omo s	apie	ns							
				(B)	CELI	. KIN	D: L	inea	r								
				(D)	CLO	IE NA	ME:	HP01	263								
		(x	i) S	EQUE	NCE	DESC	RIPI	'ION:	SEQ	ID	NO:	19:					
35						_											
																TGTCT	
	CCAC	CCCA	GC I	GGCC	CTCA	A CC	CCTC	GGCT	CTG	CTCT	CCC	GGGG	ÇTGC	AA T	GACT	CCGAT	120

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	GTGCTGAGAC	TCAACCGAGT	GAACGACGCC	CAGGAATACA	GACGGGGTGG	CCTGGGATCT	240
	CTGTTCTATC	TTACACTGGA	TGTGCTAGAG	ACTGACTGCC	ATGTGCTCAG	AAAGAAGGCA	300
	TGGCAAGACT	GTGGAATGAG	GATATTTTT	GAATCAGTTT	ATGGTCAATG	CAAAGCAATA	360
	TTTTATATGA	ACAACCCAAG	TAGAGTTCTC	TATTTAGCTG	CTTATAACTG	TACTCTTCGC	420
5	CCAGTTTCAA	AAAAAAAGAT	TTACATGACG	TGCCCTGACT	GCCCAAGCTC	CATACCCACT	480
	GACTCTTCCA	ATCACCAAGT	GCTGGAGGCT	GCCACCGAGT	CTCTTGCGAA	ATACAACAAT	540
	GAGAACACAT	CCAAGCAGTA	TTCTCTCTTC	AAAGTCACCA	GGGCTTCTAG	CCAGTGGGTG	600
	GTCGGCCCTT	CTTACTTTGT	GGAATACTTA	ATTAAAGAAT	CACCATGTAC	TAAATCCCAG	660
	GCCAGCAGCT	GTTCACTTCA	GTCCTCCGAC	TCTGTGCCTG	TTGGTCTTTG	CAAAGGTTCT	720
10	CTGACTCGAA	CACACTGGGA	AAAGTTTGTC	TCTGTGACTT	GTGACTTCTT	TGAATCACAG	780
	GCTCCAGCCA	CTGGAAGTGA	AAACTCTGCT	GTTAACCAGA	AACCTACAAA	CCTTCCCAAG	840
	GTGGAAGAAT	CCCAGCAGAA	AAACACCCCC	CCAACAGACT	CCCCTCCAA	AGCTGGGCCA	900
	AGAGGATCTG	TCCAATATCT	TCCTGACTTG	GATGATAAAA	ATTCCCAGGA	AAAGGCCCT	960
	CAGGAGGCCT	TTCCTGTGCA	TCTGGACCTA	ACCACGAATC	CCCAGGGAGA	AACCCTGGAT	1020
15	ATTTCCTTCC	TCTTCCTGGA	GCCTATGGAG	GAGAAGCTGG	TTGTCCTGCC	TTTCCCCAAA	1080
	GAAAAAGCAC	GCACTGCTGA	GTGCCCAGGG	CCAGCCCAGA	ATGCCAGCCC	TCTTGTCCTT	1140
	CCGCCA						1146

# 20 (2) INFORMATION FOR SEQ ID NO: 20:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 951
  - (B) TYPE: Nucleic acid
  - (C) STRANDEDNESS: Double
- 25 (D) TOPOLOGY: Linear
  - (ii) SEQUENCE KIND: cDNA to mRNA

## (vi) ORIGINAL SOURCE:

- (A) ORGANISM: Homo sapiens
- 30 (B) CELL KIND: Liver
  - (D) CLONE NAME: HP01299

## (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 20:

35	ATGTGGCTCT	ACCTGGCGGC	CTTCGTGGGC	CTGTACTACC	TTCTGCACTG	GTACCGGGAG	60
	AGGCAGGTGG	TGAGCCACCT	CCAAGACAAG	TATGTCTTTA	TCACGGGCTG	TGACTCGGGC	120
	TTTGGGAACC	TGCTGGCCAG	ACAGCTGGAT	GCACGAGGCT	TGAGAGTGCT	GGCTGCGTGT	180
	CTGACGGAGA	AGGGGGCCGA	GCAGCTGAGG	GGCCAGACGT	CTGACAGGCT	GGAGACGGTG	240

	ACCCTGGATG	TTACCAAGAT	GGAGAGCATC	GCTGCAGCTA	CTCAGTGGGT	GAAGGAGCAT	300
	GTGGGGGACA	GAGGACTCTG	GGGACTGGTG	AACAATGCAG	GCATTCTTAC	ACCAATTACC	360
	TTATGTGAGT	GGCTGAACAC	TGAGGACTCT	ATGAATATGC	TCAAAGTGAA	CCTCATTGGT	420
	GTGATCCAGG	TGACCTTGAG	CATGCTTCCT	TTGGTGAGGA	GAGCACGGGG	AAGAATTGTC	480
5	AATGTCTCCA	GCATTCTGGG	AAGAGTTGCT	TTCTTTGTAG	GAGGCTACTG	TGTCTCCAAG	540
	TATGGAGTGG	AAGCCTTTTC	AGATATTCTG	AGGCGTGAGA	TTCAACATTT	TGGGGTGAAA	600
	ATCAGCATAG	TTGAACCTGG	CTACTTCAGA	ACGGGAATGA	CAAACATGAC	ACAGTCCTTA	660
	GAGCGAATGA	AGCAAAGTTG	GAAAGAAGCC	CCCAAGCATA	TTAAGGAGAC	CTATGGACAG	720
	CAGTATTTTG	ATGCCCTTTA	CAATATCATG	AAGGAAGGGC	TGTTGAATTG	TAGCACAAAC	780
10	CTGAACCTGG	TCACTGACTG	CATGGAACAT	GCTCTGACAT	CGGTGCATCC	GCGAACTCGA	840
	TATTCAGCTG	GCTGGGATGC	TAAATTTTTC	TTCATCCCTC	TATCTTATTT	ACCTACATCA	900
	CTGGCAGACT	ACATTTTGAC	TAGATCTTGG	CCCAAACCAG	CCCAGGCAGT	C	951

## 15 (2) INFORMATION FOR SEQ ID NO: 21:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 888
  - (B) TYPE: Nucleic acid
  - (C) STRANDEDNESS: Double
- 20 (D) TOPOLOGY: Linear
  - (ii) SEQUENCE KIND: cDNA to mRNA

#### (vi) ORIGINAL SOURCE:

- (A) ORGANISM: Homo sapiens
- 25 (B) CELL KIND: Liver
  - (D) CLONE NAME: HP01347

## (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 21:

30	ATGAGTGACT	CCAAGGAACC	AAGGGTGCAG	CAGCTGGGCC	TCCTGGGGTG	TCTTGGCCAT	60
	GGCGCCCTGG	TGCTGCAACT	CCTCTCCTTC	ATGCTCTTGG	CTGGGGTCCT	GGTGGCCATC	120
	CTTGTCCAAG	TGTCCAAGGT	CCCCAGCTCC	CTAAGTCAGG	AACAATCCGA	GCAAGACGCA	180
	ATCTACCAGA	ACCTGACCCA	GCTTAAAGCT	GCAGTGGGTG	AGCTCTCAGA	GAAATCCAAG	240
	CTGCAGGAGA	TCTACCAGGA	GCTGACCCAG	CTGAAGGCTG	CAGTGGGTGA	GTTGCCAGAG	300
35	AAATCCAAGC	TGCAGGAGAT	CTACCAGGAG	CTGACCCGGC	TGAAGGCTGC	AGTGGGTGAG	360
	TTGCCAGAGA	AATCCAAGCT	GCAGGAGATC	TACCAGGAGC	TGACCCGGCT	GAAGGCTGCA	420
	GTGGGTGAGT	TGCCAGAGAA	ATCCAAGCTG	CAGGAGATCT	ACCAGGAGCT	GACCCGGCTG	480
	AAGGCTGCAG	TGGGTGAGTT	GCCAGAGAAA	TCCAAGCTGC	AGGAGATCTA	CCAGGAGCTG	540

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	ACGGAGCTGA	AGGCTGCAGT	GGGTGAGTTG	CCAGAGAĀAT	CCAAGCTGCA	GGAGATCTAC	600
	CAGGAGCTGA	CCCAGCTGAA	GGCTGCAGTG	GGTGAGTTGC	CAGACCAGTC	CAAGCAGCAG	660
	CAAATCTATC	AAGAACTGAC	CGATTTGAAG	ACTGCATTTG	AACGCCTGTG	CCGCCACTGT	720
	CCCAAGGACT	GGACATTCTT	CCAAGGAAAC	TGTTACTTCA	TGTCTAACTC	CCAGCGGAAC	780
5	TGGCACGACT	CCGTCACCGC	CTGCCAGGAA	GTGAGGGCCC	AGCTCGTCGT	AATCAAAACT	840
	GCTGAGGAGC	AGCTTCCAGC	GGTACTGGAA	CAGTGGAGAA	CCCAACAA		888

#### (2) INFORMATION FOR SEQ ID NO: 22:

10 (i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 591
- (B) TYPE: Nucleic acid
- (C) STRANDEDNESS: Double
- (D) TOPOLOGY: Linear
- 15 (ii) SEQUENCE KIND: cDNA to mRNA

#### (vi) ORIGINAL SOURCE:

- (A) ORGANISM: Homo sapiens
- (B) CELL KIND: Stomach cancer
- 20 (D) CLONE NAME: HP01440

#### (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 22:

	ATGTGTACGG	GAAAATGTGC	CCGCTGTGTG	GGGCTCTCCC	TCATTACCCT	CTGCCTCGTC	60
25	TGCATTGTGG	CCAACGCCCT	CCTGCTGGTA	CCTAATGGGG	AGACCTCCTG	GACCAACACC	120
	AACCATCTCA	GCTTGCAAGT	CTGGCTCATG	GGCGGCTTCA	TTGGCGGGGG	CCTAATGGTA	180
	CTGTGTCCGG	GGATTGCAGC	CGTTCGGGCA	GGGGGCAAGG	GCTGCTGTGG	TGCTGGGTGC	240
	TGTGGAAACC	GCTGCAGGAT	GCTGCGCTCG	GTCTTCTCCT	CGGCGTTCGG	GGTGCTTGGT	300
	GCCATCTACT	GCCTCTCGGT	GTCTGGAGCT	GGGCTCCGAA	ATGGACCCAG	ATGCTTAATG	360
30	AACGGCGAGT	GGGGCTACCA	CTTCGAAGAC	ACCGCGGGAG	CTTACTTGCT	CAACCGCACT	420
	CTATGGGATC	GGTGCGAGGC	GCCCCTCGC	GTGGTCCCCT	GGAATGTGAC	GCTCTTCTCG	480
	CTGCTGGTGG	CCGCCTCCTG	CCTGGAGATA	GTACTGTGTG	GGATCCAGCT	GGTGAACGCG	540
	ACCATTGGTG	TCTTCTGCGG	CGATTGCAGG	AAAAAACAGG	ACACCCCTCA	С	591

- (2) INFORMATION FOR SEQ ID NO: 23:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 663

		109		
		(B) TYPE: Nucleic acid		
		(C) STRANDEDNESS: Double		
		(D) TOPOLOGY: Linear		
	(ii)	SEQUENCE KIND: cDNA to mRNA		
5				
	(vi)	ORIGINAL SOURCE:		
		(A) ORGANISM: Homo sapiens		
		(B) CELL KIND: Stomach cancer		
		(D) CLONE NAME: HP01526		
10				
	(xi)	SEQUENCE DESCRIPTION: SEQ ID NO: 23:		
•				
	ATGGAGGCGG	GCGGCTTTCT GGACTCGCTC ATTTACGGAG CATGCG	TGGT CTTCACCCTT	60
	GGCATGTTCT	CCGCCGGCCT CTCGGACCTC AGGCACATGC GAATGAC	CCCG GAGTGTGGAC	120
15	AACGTCCAGT	TCCTGCCCTT TCTCACCACG GAAGTCAACA ACCTGG	GCTG GCTGAGTTAT	180
	GGGGCTTTGA	AGGGAGACGG GATCCTCATC GTCGTCAACA CAGTGGC	GTGC TGCGCTTCAG	240
	ACCCTGTATA	TCTTGGCATA TCTGCATTAC TGCCCTCGGA AGCGTG	TTGT GCTCCTACAG	300
	ACTGCAACCC	TGCTAGGGGT CCTTCTCCTG GGTTATGGCT ACTTTTC	GGCT CCTGGTACCC	360
	AACCCTGAGG	CCCGGCTTCA GCAGTTGGGC CTCTTCTGCA GTGTCTT	TCAC CATCAGCATG	420
20		CACTGGCTGA CTTGGCTAAG GTGATTCAAA CTAAATC		480
		TCACCATTGC TACCCTTCTC ACCTCTGCCT CCTGGTG		540
		ATCCCTATAT CATGGTGTCC AACTTTCCAG GAATCGT		600
	CGCTTCTGGC	TTTTCTGGAA GTACCCCCAG GAGCAAGACA GGAACTA	ACTG GCTCCTGCAA	660
	ACC			663
25				
		TION FOR SEQ ID NO: 24:		
	(i) S	EQUENCE CHARACTERISTICS:		
	•	(A) LENGTH: 753		
30		(B) TYPE: Nucleic acid		
		(C) STRANDEDNESS: Double	•	
		(D) TOPOLOGY: Linear		
	(ii)	SEQUENCE KIND: cDNA to mRNA		
35	/1 5	ORIGINAL COURGE		
<i>.</i> .	(V1)	ORIGINAL SOURCE:		
		(A) ORGANISM: Homo sapiens  (B) CELL KIND: Stomach cancer		
		COL SUBLIS REDUCE STOMPED CONCAT		

(D) CLONE NAME: HP10230

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#### (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 24:

	ATGTCGGACA	TCGGAGACTG	GTTCAGGAGC	ATCCCGGCGA	TCACGCGCTA	TTGGTTCGCC	60
	GCCACCGTCG	CCGTGCCCTT	GGTCGGCAAA	CTCGGCCTCA	TCAGCCCGGC	CTACCTCTTC	120
5	CTCTGGCCCG	AAGCCTTCCT	TTATCGCTTT	CAGATTTGGA	GGCCAATCAC	TGCCACCTTT	180
	TATTTCCCTG	TGGGTCCAGG	AACTGGATTT	CTTTATTTGG	TCAATTTATA	TTTCTTATAT	240
	CAGTATTCTA	CGCGACTTGA	AACAGGAGCT	TTTGATGGGA	GGCCAGCAGA	CTATTTATTC	300
	ATGCTCCTCT	TTAACTGGAT	TTGCATCGTG	ATTACTGGCT	TAGCAATGGA	TATGCAGTTG	360
	CTGATGATTC	CTCTGATCAT	GTCAGTACTT	TATGTCTGGG	CCCAGCTGAA	CAGAGACATG	420
10	ATTGTATCAT	TTTGGTTTGG	AACACGATTT	AAGGCCTGCT	ATTTACCCTG	GGTTATCCTT	480
	GGATTCAACT	ATATCATCGG	AGGCTCGGTA	ATCAATGAGC	TTATTGGAAA	TCTGGTTGGA	540
	CATCTTTATT	TTTTCCTAAT	GTTCAGATAC	CCAATGGACT	TGGGAGGAAG	AAATTTTCTA	600
	TCCACACCTC	AGTTTTTGTA	CCGCTGGCTG	CCCAGTAGGA	GAGGAGGAGT	ATCAGGATTT	660
	GGTGTGCCCC	CTGCTAGCAT	GAGGCGAGCT	GCTGATCAGA	ATGGCGGAGG	CGGGAGACAC	720
15	AACTGGGGCC	AGGGCTTTCG	ACTTGGAGAC	CAG		•	753

#### (2) INFORMATION FOR SEQ ID NO: 25:

(i) SEQUENCE CHARACTERISTICS:

20

- (A) LENGTH: 318
- (B) TYPE: Nucleic acid
- (C) STRANDEDNESS: Double
- (D) TOPOLOGY: Linear
- (ii) SEQUENCE KIND: cDNA to mRNA

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- (vi) ORIGINAL SOURCE:
  - (A) ORGANISM: Homo sapiens
  - (B) CELL KIND: Epidermoid carcinoma
  - (C) CELL LINE: KB

30

(D) CLONE NAME: HP10389

#### (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 25:

	ATGGCGACTC	CCGGCCCTGT	GATTCCGGAG	GTCCCCTTTG	AACCATCGAA	GCCTCCAGTC	60
35	ATTGAGGGGC	TGAGCCCCAC	TGTTTACAGG	AATCCAGAGA	GTTTCAAGGA	AAAGTTCGTT	120
	CGCAAGACCC	GCGAGAACCC	GGTGGTACCC	ATAGGTTGCC	TGGCCACGGC	GGCCGCCCTC	180
	ACCTACGGCC	TCTACTCCTT	CCACCGGGGC	AACAGCCAGC	GCTCTCAGCT	CATGATGCGC	240
	ACCCGGATCG	CCGCCCAGGG	TTTCACGGTC	GCAGCCATCT	TGCTGGGTCT	GGCTGTCACT	300

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	GCTATGAAGT CTCGACCC	318
	(2) INFORMATION FOR SEQ ID NO: 26:	
5	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 234	
	(B) TYPE: Nucleic acid	
	(C) STRANDEDNESS: Double	
	(D) TOPOLOGY: Linear	
10	(ii) SEQUENCE KIND: cDNA to mRNA	
	(vi) ORIGINAL SOURCE:	
	(A) ORGANISM: Homo sapiens	
	(B) CELL KIND: Stomach cancer	
15	(D) CLONE NAME: HP10408	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 26:	
	ATGGGGTCTG GGCTGCCCCT TGTCCTCCTC TTGACCCTCC TTGGCAGCTC ACATGGAACA	60
20	GGGCCGGGTA TGACTTTGCA ACTGAAGCTG AAGGAGTCTT TTCTGACAAA TTCCTCCTAT	120
	GAGTCCAGCT TCCTGGAATT GCTTGAAAAG CTCTGCCTCC TCCTCCATCT CCCTTCAGGG	180
	ACCAGCGTCA CCCTCCACCA TGCAAGATCT CAACACCATG TTGTCTGCAA CACA	234
25	(C) THEORYAMION DOD ODG TO NO. OT	
25	(2) INFORMATION FOR SEQ ID NO: 27:	
	(1) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 942	
	(B) TYPE: Nucleic acid (C) STRANDEDNESS: Double	
30	(D) TOPOLOGY: Linear	
50	(ii) SEQUENCE KIND: cDNA to mRNA	
	(vi) ORIGINAL SOURCE:	
	(A) ORGANISM: Homo sapiens	
35	(B) CELL KIND: Stomach cancer	
-	, , <del>value .</del>	

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 27:

(D) CLONE NAME: HP10412

	ATGGTGGCGC	CTGTGTGGTA	CTTGGTAGCG	GCGGCTCTGC	TAGTCGGCTT	TATCCTCTTC	6
*	CTGACTCGCA	GCCGGGGCCG	GGCGGCATCA	GCCGGCCAAG	AGCCACTGCA	CAATGAGGAG	120
	CTGGCAGGAG	CAGGCCGGGT	GGCCCAGCCT	GGGCCCCTGG	AGCCTGAGGA	GCCGAGAGCT	180
•	GGAGGCAGGC	CTCGGCGCCG	GAGGGACCTG	GGCAGCCGCC	TACAGGCCCA	GCGTCGAGCC	240
5	CAGCGGGTGG	CCTGGGCAGA	AGCAGATGAG	AACGAGGAGG	AAGCTGTCAT	CCTAGCCCAG	300
	GAGGAGGAAG	GTGTCGAGAA	GCCAGCGGAA	ACTCACCTGT	CGGGGAAAAT	TGGAGCTAAG	360
	AAACTGCGGA	AGCTGGAGGA	GAAACAAGCG	CGAAAGGCCC	AGCGTGAGGC	AGAGGAGGCT	420
	GAACGTGAGG	AGCGGAAACG	ACTCGAGTCC	CAGCGCGAAG	CTGAGTGGAA	GAAGGAGGAG	480
	GAGCGGCTTC	GCCTGGAGGA	GGAGCAGAAG	GAGGAGGAGG	AGAGGAAGGC	CCGCGAGGAG	540
10	CAGGCCCAGC	GGGAGCATGA	GGAGTACCTG	AAACTGAAGG	AGGCCTTTGT	GGTGGAGGAG	600
	GAAGGCGTAG	GAGAGACCAT	GACTGAGGAA	CAGTCCCAGA	GCTTCCTGAC	AGAGTTCATC	660
	AACTACATCA	AGCAGTCCAA	GGTTGTGCTC	TTGGAAGACC	TGGCTTCCCA	GGTGGGCCTA	720
	CGCACTCAGG	ACACCATAAA	TCGCATCCAG	GACCTGCTGG	CTGAGGGGAC	TATAACAGGT	780
	GTGATTGACG	ACCGGGGCAA	GTTCATCTAC	ATAACCCCAG	AGGAACTGGC	CGCCGTGGCC	840
L5	AACTTCATCC	GACAGCGGGG	CCGGGTGTCC	ATCGCCGAGC	TTGCCCAAGC	CAGCAACTCC	900
	CTCATCGCCT	GGGGCCGGGA	GTCCCCTGCC	CAAGCCCCAG	CC		942

#### (2) INFORMATION FOR SEQ ID NO: 28:

20 (i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 585
- (B) TYPE: Nucleic acid
- (C) STRANDEDNESS: Double
- (D) TOPOLOGY: Linear
- 25 (ii) SEQUENCE KIND: cDNA to mRNA

#### (vi) ORIGINAL SOURCE:

- (A) ORGANISM: Homo sapiens
- (B) CELL KIND: Stomach cancer
- 30 (D) CLONE NAME: HP10413

#### (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 28:

	ATGGCTGCCG	AGGATGTGGT	GGCGACTGGC	GCCGACCCAA	GCGATCTGGA	GAGCGGCGGG	60
35	CTGCTGCATG	AGATTTTCAC	GTCGCCGCTC	AACCTGCTGC	TGCTTGGCCT	CTGCATCTTC	120
	CTGCTCTACA	AGATCGTGCG	CGGGGACCAG	CCGGCGGCCA	GCGGCGACAG	CGACGACGAC	180
	GAGCCGCCCC	CTCTGCCCCG	CCTCAAGCGG	CGCGACTTCA	CCCCGCCGA	GCTGCGGCGC	240
	TTCGACGCC	TCCAGGACCC	GCGCATACTC	ATGGCCATCA	ACGGCAAGGT	GTTCGATGTG	300

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	ACCAAAGGCC	GCAAATTCTA	CGGGCCCGAG	GGGCCGTATG	GGGTCTTTGC	TGGAAGAGAT	360
	GCATCCAGGG	GCCTTGCCAC	ATTTTGCCTG	GATAAGGAAG	CACTGAAGGA	TGAGTACGAT	420
	GACCTTTCTG	ACCTCACTGC	TGCCCAGCAG	GAGACTCTGA	GTGACTGGGA	GTCTCAGTTC	480
	ACTTTCAAGT	ATCATCACGT	GGGCAAACTG	CTGAAGGAGG	GGGAGGAGCC	CACTGTGTAC	540
5	TCAGATGAGG	AAGAACCAAA	AGATGAGAGT	GCCCGGAAAA	ATGAT		585

#### (2) INFORMATION FOR SEQ ID NO: 29:

(i) SEQUENCE CHARACTERISTICS:

10

- (A) LENGTH: 1386
- (B) TYPE: Nucleic acid
- (C) STRANDEDNESS: Double
- (D) TOPOLOGY: Linear
- (ii) SEQUENCE KIND: cDNA to mRNA

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#### (vi) ORIGINAL SOURCE:

- (A) ORGANISM: Homo sapiens
- (B) CELL KIND: Stomach cancer
- (D) CLONE NAME: HP10415

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#### (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 29:

	ATGTTGGACT	TCGCGATCTT	CGCCGTTACC	TTCTTGCTGG	CGTTGGTGGG	AGCCGTGCTC	. 60
	TACCTCTATC	CGGCTTCCAG	ACAAGCTGCA	GGAATTCCAG	GGATTACTCC	AACTGAAGAA	120
25	AAAGATGGTA	ATCTTCCAGA	TATTGTGAAT	AGTGGAAGTT	TGCATGAGTT	CCTGGTTAAT	180
	TTGCATGAGA	GATATGGGCC	TGTGGTCTCC	TTCTGGTTTG	GCAGGCGCCT	CGTGGTTAGT	240
	TTGGGCACTG	TTGATGTACT	GAAGCAGCAT	ATCAATCCCA	ATAAGACATT	GGACCCTTTT	300
	GAAACCATGC	TGAAGTCATT	ATTAAGGTAT	CAATCTGGTG	GTGGCAGTGT	GAGTGAAAAC	360
	CACATGAGGA	AAAAATTGTA	TGAAAATGGT	GTGACTGATT	CTCTGAAGAG	TAACTTTGCC	420
30	CTCCTCCTAA	AGCTTTCAGA	AGAATTATTA	GATAAATGGC	TCTCCTACCC	AGAGACCCAG	480
	CACGTGCCCC	TCAGCCAGCA	TATGCTTGGT	TTTGCTATGA	AGTCTGTTAC	ACAGATGGTA	540
	ATGGGTAGTA	CATTTGAAGA	TGATCAGGAA	GTCATTCGCT	TCCAGAAGAA	TCATGGCACA	600
	GTTTGGTCTG	AGATTGGAAA	AGGCTTTCTA	GATGGGTCAC	TTGATAAAAA	CATGACTCGG	660
	AAAAAACAAT	ATGAAGATGC	CCTCATGCAA	CTGGAGTCTG	TTTTAAGGAA	CATCATAAAA	720
35	GAACGAAAAG	GAAGGAACTT	CAGTCAACAT	ATTTTCATTG	ACTCCTTAGT	ACAAGGGAAC	780
	CTTAATGACC	AACAGATCCT	AGAAGACAGT	ATGATATTTT	CTCTGGCCAG	TTGCATAATA	840
	ACTGCAAAAT	TGTGTACCTG	GGCAATCTGT	TTTTTAACCA	CCTCTGAAGA	AGTTCAAAAA	900
	AAATTATATG	AAGAGATAAA	CCAAGTTTTT	GGAAATGGTC	CTGTTACTCC	AGAGAAAATT	960

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	GAGCAGCTCA	GATATTGTCA	GCATGTGCTT	TGTGAAACTG	TTCGAACTGC	CAAACTGACT	1020
	CCAGTTTCTG	CCCAGCTTCA	AGATATTGAA	GGAAAAATTG	ACCGATTTAT	TATTCCTAGA	1080
	GAGACCCTCG	TCCTTTATGC	CCTTGGTGTG	GTACTTCAGG	ATCCTAATAC	TTGGCCATCT	1140
	CCACACAAGT	TTGATCCAGA	TCGGTTTGAT	GATGAATTAG	TAATGAAAAC	TTTTTCCTCA	1200
5	CTTGGATTCT	CAGGCACACA	GGAGTGTCCA	GAGTTGAGGT	TTGCATATAT	GGTGACCACA	1260
	GTACTTCTTA	GTGTATTGGT	GAAGAGACTG	CACCTACTTT	CTGTGGAGGG	ACAGGTTATT	1320
	GAAACAAAGT	ATGAACTGGT	AACATCATCA	AGGGAAGAAG	CTTGGATCAC	TGTCTCAAAG	1380
	AGATAT						1386
0							
	(2) THEODMA	מס מסע ואחדיי	o in Mo. 20	١.			

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- (2) INFORMATION FOR SEQ ID NO: 30:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 741
    - (B) TYPE: Nucleic acid
- 15 (C) STRANDEDNESS: Double
  - (D) TOPOLOGY: Linear
  - (ii) SEQUENCE KIND: cDNA to mRNA

#### (vi) ORIGINAL SOURCE:

20

- (A) ORGANISM: Homo sapiens
- (B) CELL KIND: Stomach cancer
- (D) CLONE NAME: HP10419

#### (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 30:

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30

ATGGGGGCTG	CGGTGTTTTT	CGGCTGCACT	TTCGTCGCGT	TCGGCCCGGC	CTTCGCGCTT	60
TTCTTGATCA	CTGTGGCTGG	GGACCCGCTT	CGCGTTATCA	TCCTGGTCGC	AGGGGCATTT	120
TTCTGGCTGG	TCTCCCTGCT	CCTGGCCTCT	GTGGTCTGGT	TCATCTTGGT	CCATGTGACC	180
GACCGGTCAG	ATGCCCGGCT	CCAGTACGGC	CTCCTGATTT	TTGGTGCTGC	TGTCTCTGTC	240
CTTCTACAGG	AGGTGTTCCG	CTTTGCCTAC	TACAAGCTGC	TTAAGAAGGC	AGATGAGGGG	300
TTAGCATCGC	TGAGTGAGGA	CGGAAGATCA	CCCATCTCCA	TCCGCCAGAT	GGCCTATGTT	360
TCTGGTCTCT	CCTTCGGTAT	CATCAGTGGT	GTCTTCTCTG	TTATCAATAT	TTTGGCTGAT	420
GCACTTGGGC	CAGGTGTGGT	TGGGATCCAT	GGAGACTCAC	CCTATTACTT	CCTGACTTCA	480
GCCTTTCTGA	CAGCAGCCAT	TATCCTGCTC	CATACCTTTT	GGGGAGTTGT	GTTCTTTGAT	540
GCCTGTGAGA	GGAGACGGTA	CTGGGCTTTG	GGCCTGGTGG	TTGGGAGTCA	CCTACTGACA	600
TCGGGACTGA	CATTCCTGAA	CCCCTGGTAT	GAGGCGAGCC	TGCTGCCCAT	CTATGCAGTC	660
ACTGTTTCCA	TGGGGCTCTG	GGCCTTCATC	ACAGCTGGAG	GGTCCCTCCG	AAGTATTCAG	720
CGCAGCCTCT	TGTGTAAGGA	С	•			741

	(2) INFORMATION FOR SEQ ID NO: 31:	
	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 339	
5	(B) TYPE: Nucleic acid	
	(C) STRANDEDNESS: Double	
	(D) TOPOLOGY: Linear	
	(ii) SEQUENCE KIND: cDNA to mRNA	
10	(vi) ORIGINAL SOURCE:	
	(A) ORGANISM: Homo sapiens	
•	(B) CELL KIND: Stomach cancer	
	(D) CLONE NAME: HP10424	
15	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 31:	
	ATGAACTTCT ATTTACTCCT AGCGAGCAGC ATTCTGTGTG CCTTGATTGT CTTCTGGAAA	60
	TATCGCCGCT TTCAGAGAAA CACTGGCGAA ATGTCATCAA ATTCAACTGC TCTTGCACTA	120
20	GTGAGACCCT CTTCTTCTGG GTTAATTAAC AGCAATACAG ACAACAATCT TGCAGTCTAC	180
20	GACCTCTCTC GGGATATTTT AAATAATTTC CCACACTCAA TAGCCAGGCA GAAGCGAATA	240
	TTGGTAAACC TCAGTATGGT GGAAAACAAG CTGGTTGAAC TGGAACATAC TCTACTTAGC	300
	AAGGGTTTCA GAGGTGCATC ACCTCACCGG AAATCCACC	339
25	(2) INFORMATION FOR SEQ ID NO: 32:	
	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 1095	
	(B) TYPE: Nucleic acid	
	(C) STRANDEDNESS: Double	
30	(D) TOPOLOGY: Linear	
	(ii) SEQUENCE KIND: cDNA to mRNA	
٠	(vi) ORIGINAL SOURCE:	
	(A) ORGANISM: Homo saniens	

(B) CELL KIND: Epidermoid carcinoma

(C) CELL LINE: KB

(D) CLONE NAME: HP10428

## (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 32:

	ATGGGGAGGT	GGGCCCTCGA	TGTGGCCTTT	TTGTGGAAGG	CGGTGTTGAC	CCTGGGGCTG	60
	GTGCTTCTCT	ACTACTGCTT	CTCCATCGGC	ATCACCTTCT	ACAACAAGTG	GCTGACAAAG	120
5	AGCTTCCATT	TCCCCCTCTT	CATGACGATG	CTGCACCTGG	CCGTGATCTT	CCTCTTCTCC	180
	GCCCTGTCCA	GGGCGCTGGT	TCAGTGCTCC	AGCCACAGGG	CCCGTGTGGT	GCTGAGCTGG	240
	GCCGACTACC	TCAGAAGAGT	GGCTCCCACA	GCTCTGGCGA	CGGCGCTTGA	CGTGGGCTTG	300
	TCCAACTGGA	GCTTCCTGTA	TGTCACCGTC	TCGCTGTACA	CAATGACCAA	ATCCTCAGCT	360
	GTCCTCTTCA	TCTTGATCTT	CTCTCTGATC	TTCAAGCTGG	AGGAGCTGCG	CGCGGCACTG	420
10	GTCCTGGTGG	TCCTCCTCAT	CGCCGGGGGT	CTCTTCATGT	TCACCTACAA	GTCCACACAG	480
	TTCAACGTGG	AGGGCTTCGC	CTTGGTGCTG	GGGGCCTCGT	TCATCGGTGG	CATTCGCTGG	540
•	ACCCTCACCC	AGATGCTCCT	GCAGAAGGCT	GAACTCGGCC	TCCAGAATCC	CATCGACACC	600
	ATGTTCCACC	TGCAGCCACT	CATGTTCCTG	GGGCTCTTCC	CTCTCTTTGC	TGTATTTGAA	660
	GGTCTCCATT	TGTCCACATC	TGAGAAAATC	TTCCGTTTCC	AGGACACAGG	GCTGCTCCTG	720
15	CGGGTACTTG	GGAGCCTCTT	CCTTGGCGGG	ATTCTCGCCT	TTGGTTTGGG	CTTCTCTGAG	780
	TTCCTCCTGG	TCTCCAGAAC	CTCCAGCCTC	ACTCTCTCCA	TTGCCGGCAT	TTTTAAGGAA	840
	GTCTGCACTT	TGCTGTTGGC	AGCTCATCTG	CTGGGCGATC	AGATCAGCCT	CCTGAACTGG	900
	CTGGGCTTCG	CCCTCTGCCT	CTCGGGAATA	TCCCTCCACG	TTGCCCTCAA	AGCCCTGCAT	960
	TCCAGAGGTG	ATGGTGGCCC	CAAGGCCTTG	AAGGGGCTGG	GCTCCAGCCC	CGACCTGGAG	1020
20	CTGCTGCTCC	GGAGCAGCCA	GCGGGAGGAA	GGTGACAATG	AGGAGGAGGA	GTACTTTGTG	1080
	GCCCAGGGGC	AGCAG					1095

## (2) INFORMATION FOR SEQ ID NO: 33:

- 25 (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 678
  - (B) TYPE: Nucleic acid
  - (C) STRANDEDNESS: Double
  - (D) TOPOLOGY: Linear
- 30 (ii) SEQUENCE KIND: cDNA to mRNA
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo sapiens
    - (B) CELL KIND: Stomach cancer
- 35 (D) CLONE NAME: HP10429
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 33:

117

	117	
	ATGCCTACCA CAAAGAAGAC ATTGATGTTC TTATCAAGCT TTTTCACCAG CCTTGGGTCC	60
	TTCATTGTAA TTTGCTCTAT TCTTGGGACA CAAGCATGGA TCACCAGTAC AATTGCTGTT	120
	AGAGACTCTG CTTCAAATGG GAGCATTTTC ATCACTTACG GACTTTTTCG TGGGGAGAGT	180
	AGTGAAGAAT TGAGTCACGG ACTTGCAGAA CCAAAGAAAA AGTTTGCAGT TTTAGAGATA	240
5	CTGAATAATT CTTCCCAAAA AACTCTGCAT TCGGTGACTA TCCTGTTCCT GGTCCTGAGT	300
	TTGATCACGT CGCTGCTGAG CTCTGGGTTT ACCTTCTACA ACAGCATCAG CAACCCTTAC	360
	CAGACATTCC TGGGGCCGAC GGGGGTGTAC ACCTGGAACG GGCTCGGTGC ATCCTTCGTT	420
	TTTGTGACCA TGATACTGTT TGTGGCGAAC ACGCAGTCCA ACCAACTCTC CGAAGAGTTG	480
	TTCCAAATGC TTTACCCGGC AACCACCAGT AAAGGAACGA CCCACAGTTA CGGATACTCG	540
10	TTCTGGCTCA TACTGCTCGT CATTCTTCTA AATATAGTCA CTGTAACCAT CATCATTTTC	600
	TACCAGAAGG CCAGATACCA GCGGAAGCAG GAGCAGAGAA AGCCAATGGA ATATGCTCCA	660
	AGGGACGGAA TTTTATTC	678
15	(2) INFORMATION FOR SEQ ID NO: 34:	
	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 387	
	(B) TYPE: Nucleic acid	
	(C) STRANDEDNESS: Double	
20	(D) TOPOLOGY: Linear	
	(ii) SEQUENCE KIND: cDNA to mRNA	
	(vi) ORIGINAL SOURCE:	
	(A) ORGANISM: Homo sapiens	
25	(B) CELL KIND: Liver	
	(D) CLONE NAME: HP10432	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 34:	
30		
	ATGGCTCGGG GCTCGCTGCG CCGGTTGCTG CGGCTCCTCG TGCTGGGGCT CTGGCTGG	60
	TTGCTGCGCT CCGTGGCCGG GGAGCAAGCG CCAGGCACCG CCCCCTGCTC CCGCGGCAGC	120
	TCCTGGAGCG CGGACCTGGA CAAGTGCATG GACTGCGCGT CTTGCAGGGC GCGACCGCAC	180
	AGCGACTTCT GCCTGGGCTG CGCTGCAGCA CCTCCTGCCC CCTTCCGGCT GCTTTGGCCC	240
35	ATCCTTGGGG GCGCTCTGAG CCTGACCTTC GTGCTGGGGC TGCTTTCTGG CTTTTTGGTC	300
	TGGAGACGAT GCCGCAGGAG AGAGAAGTTC ACCACCCCCA TAGAGGAGAC CGGCGGAGAG	360

387

GGCTGCCCAG CTGTGGCGCT GATCCAG

WO 98/55508

	(2) INFORMATION FOR SEQ ID NO: 35:	
	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 489	
	(B) TYPE: Nucleic acid	
5	(C) STRANDEDNESS: Double	
	(D) TOPOLOGY: Linear	
	(ii) SEQUENCE KIND: cDNA to mRNA	
	(vi) ORIGINAL SOURCE:	
10	(A) ORGANISM: Homo sapiens	
	(B) CELL KIND: Liver	
•	(D) CLONE NAME: HP10433	
<u>.</u> _	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 35:	
15		
	ATGCGACGGC TGCTGATCCC TCTGGCCCTG TGGCTGGGCG CGGTGGGCGT GGGCGTCGCC	60
	GAGCTCACGG AAGCCCAGCG CCGGGGCCTG CAGGTGGCCC TGGAGGAATT TCACAAGCAC	120
	CCGCCCGTGC AGTGGGCCTT CCAGGAGACC AGTGTGGAGA GCGCCGTGGA CACGCCCTTC	180
20	CCAGCTGGAA TATTTGTGAG GCTGGAATTT AAGCTGCAGC AGACAAGCTG CCGGAAGAGG	240
20	GACTGGAAGA AACCCGAGTG CAAAGTCAGG CCCAATGGGA GGAAACGGAA ATGCCTGGCC	300
	TGCATCAAAC TGGGCTCTGA GGACAAAGTT CTGGGCCGGT TGGTCCACTG CCCCATAGAG	360
	ACCCAAGTTC TGCGGGAGGC TGAGGAGCCC CAGGAGACCC AGTGCCTCAG GGTGCAGCGG	420
	GCTGGTGAGG ACCCCCACAG CTTCTACTTC CCTGGACAGT TCGCCTTCTC CAAGGCCCTG CCCCGCAGC	480
25	COCCOCAGO	489
	(2) INFORMATION FOR SEQ ID NO: 36:	
	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 579	
30	(B) TYPE: Nucleic acid	
	(C) STRANDEDNESS: Double	
	(D) TOPOLOGY: Linear	
	(ii) SEQUENCE KIND: cDNA to mRNA	
35	(vi) ORIGINAL SOURCE:	
	(A) ORGANISM: Homo sapiens	
	(B) CELL KIND: Stomach cancer	

(D) CLONE NAME: HP10480

## (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 36:

	ATGATCCGCT GCGGCCTGGC CTGCGAGCGC TGCCGCTGGA TCCTGCCCCT GCTCCTACTC	6
	AGCGCCATCG CCTTCGACAT CATCGCGCTG GCCGGCCGCG GCTGGTTGCA GTCTAGCGAC	12
5	CACGGCCAGA CGTCCTCGCT GTGGTGGAAA TGCTCCCAAG AGGGCGGCGG CAGCGGGTCC	18
	TACGAGGAGG GCTGTCAGAG CCTCATGGAG TACGCGTGGG GTAGAGCAGC GGCTGCCATG	24
	CTCTTCTGTG GCTTCATCAT CCTGGTGATC TGTTTCATCC TCTCCTTCTT CGCCCTCTGT	30
	GGACCCCAGA TGCTTGTCTT CCTGAGAGTG ATTGGAGGTC TCCTTGCCTT GGCTGCTGTG	36
	TTCCAGATCA TCTCCCTGGT AATTTACCCC GTGAAGTACA CCCAGACCTT CACCCTTCAT	42
10	GCCAACCGTG CTGTCACTTA CATCTATAAC TGGGCCTACG GCTTTGGGTG GGCAGCCACG	48
	ATTATCCTGA TCGGCTGTGC CTTCTTCTTC TGCTGCCTCC CCAACTACGA AGATGACCTT	54
	CTGGGCAATG CCAAGCCCAG GTACTTCTAC ACATCTGCC	57
15	(2) INFORMATION FOR SEQ ID NO: 37:	
	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 1502	
	(B) TYPE: Nucleic acid	
	(C) STRANDEDNESS: Double	
20	(D) TOPOLOGY: Linear	
	(ii) SEQUENCE KIND: cDNA to mRNA	
	(with one of the course	
	(vi) ORIGINAL SOURCE:	
25	(A) ORGANISM: Homo sapiens	
23	(B) CELL KIND: Liver	
	(D) CLONE NAME: HP01263	
	(ix) SEQUENCE CHARACTERISTICS:	
	(A) CHARACTERIZATION CODE: CDS	
30	(B) EXISTENCE POSITION: 37 1185	
	(C) CHARACTERIZATION METHOD: E	
	· · · · · · · · · · · · · · · · · · ·	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 37:	
	·	
35	ACAAACTGAC CCATCCTGGG CCTTGTTCTC CACAGA ATG GGT CTG CTC CTT CCC	54
	Met Gly Leu Leu Pro	
	1 5	
	CTC CCA CTC TCC ATC CTA CTC CTC TCC TCC	100

	Leu	ı Ala	Lev	Cys	Ile	Leu	Val	Leu	Cys	Cys	Gly	Als	Met	. Sei	Pro	Pro	
				10	)				15	,				20	)		
	CAG	CTG	GCC	CTC	AAC	CCC	TCG	GCT	CTG	CTC	TCC	CGG	GGC	TGC	TAA C	GAC	150
	G1n	Leu	Ala	Leu	Asn	Pro	Ser	Ala	Leu	Leu	Ser	Arg	Gly	Cys	Asn	Asp	
5			25					30					35	ı			
																AAA	198
	Ser			Leu	Ala	Val	Ala	Gly	Phe	Ala	Leu	Arg	Asp	Ile	Asn	Lys	
		40					45					50					
															GAC		246
10			Lys	Asp	Gly		Va1	Leu	Arg	Leu	Asn	Arg	Val	Asn	Asp	Ala	
	55					60					65					70	
															ACA		294
	Gin	GIU	Tyr	Arg		Gly	Gly	Leu	Gly		Leu	Phe	Tyr	Leu	Thr	Leu	
1 E	C 4 m	C M C	C 111 4	040	75	212				80					85		
15															TGG		342
	Asp	Val	ren		Inr	Asp	Cys	His		Leu	Arg	Lys	Lys		Trp	Gln	
	CAC	መድመ	CCA	90	400	4 T A	mmm		95	<b>m</b> o.				100			
					*	•									TGC		390
20	vsh	Cys	105	met	ALG	116	rne		GIU	Ser	Val	Tyr		Gin	Cys	ràs	
20	GCA	АТА		ጥልጥ	A ጥር	AAC	AAC	110	ልርጥ	ACA	C TO TO	CTC	115	ጥጥለ	GCT	CCT	438
	•														Ala		430
		120		-,-			125		502	6	,41	130	-7-	Dou		*****	
	TAT		TGT	ACT	CTT	CGC		GTT	TCA	AAA	AAA		АТТ	TAC	ATG	ACG	486
25															Met		,,,,
	135		·			140				-,-	145	-,-		- , -		150	
	TGC	CCT	GAC	TGC	CCA	AGC	TCC	ATA	CCC	ACT	GAC	TCT	TCC	AAT	CAC	CAA	534
															His		
					155					160	•				165		
30	GTG	CTG	GAG	GCT	GCC	ACC	GAG	TCT	CTT	GCG	AAA	TAC	AAC	AAT	GAG	AAC	582
	Val	Leu	Glu	Ala	Ala	Thr	Glu	Ser	Leu	Ala	Lys	Tyr	Asn	Asn	Glu	Asn	
				170					175					180			
	ACA	TCC	AAG	CAG	TAT	TCT	CTC	TTC	AAA	GTC	ACC	AGG	GCT	TCT	AGC	CAG	630
	Thr	Ser	Lys	Gln	Tyr	Ser	Leu	Phe	Lys	Va1	Thr	Arg	Ala	Ser	Ser	Gln	
35			185					190					195				
	TGG	GTG	GTC	GGC	CCT	TCT	TAC	TTT	GTG	GAA	TAC	TTA	ATT	AAA	GAA	TCA	678
	Trp	Val	Val	Gly	Pro	Ser	Tyr	Phe	Val	Glu	Tyr	Leu	Ile	Lys	Glu	Ser	
		200					205					210					

	CCA	TGT	ACT	AAA	TCC	CAG	GCC	AGC	AGC	TGT	TCA	CTT	CAG	TCC	TCC	GAC	726
	Pro	Cys	Thr	Lys	Ser	Gln	Ala	Ser	Ser	Cys	Ser	Leu	Gln	Ser	Ser	Asp	
	215					220					225					230	
	TCT	GTG	CCT	GTT	GGT	CTT	TGC	AAA	GGT	TCT	CTG	ACT	CGA	ACA	CAC	TGG	774
5	Ser	Val	Pro	Va1	Gly	Leu	Cys	Lys	G1y	Ser	Leu	Thr	Arg	Thr	His	Trp	
					235					240					245		
	GAA	AAG	TTT	GTC	TCT	GTG	ACT	TGT	GAC	TTC	TTT	GAA	TCA	CAG	GCT	CCA	822
			Phe														
•				250				·	255					260			
10	GCC	ACT	GGA	AGT	GAA	AAC	TCT	GCT	GTT	AAC	CAG	AAA	CCT	ACA	AAC	CTT	870
			Gly														
			265					270				•	275				
	ccc	AAG	GTG	GAA	GAA	TCC	CAG	CAG	AAA	AAC	ACC	CCC	CCA	ACA	GAC	TCC	918
			Val														
15		280					285					290			•		
	ccc	TCC	AAA	GCT	GGG	CCA	AGA	GGA	TCT	GTC	CAA	TAT	CTT	CCT	GAC	TTG	966
			Lys														
	295					300	_	-			305	•			•	310	
	GAT	GAT	AAA	AAT	TCC	CAG	GAA	AAG	GGC	CCT	CAG	GAG	GCC	TTT	ССТ	GTG	1014
20			Lys														
					315			•	·	320					325		
	CAT	CTG	GAC	CTA	ACC	ACG	TAA	ccc	CAG	GGA	GAA	ACC	CTG	GAT	ATT	TCC	1062
			Asp		_												
				330					335	•				340			
25	TTC	CTC	TTC	CTG	GAG	CCT	ATG	GAG	GAG	AAG	CTG	GTT	GTC	CTG	CCT	TTC	1110
			Phe														
			345					350					355				
	ccc	AAA	GAA	AAA	GCA	CGC	ACT	GCT	GAG	TGC	CCA	GGG	CCA	GCC	CAG .	AAT	1158
	Pro	Lys	Glu	Lys	Ala	Arg	Thr	Ala	Glu	Cys	Pro	Gly	Pro	Ala	Gln .	Asn	
30		360					365					370					
	GCC	AGC	CCT	CTT	GTC	CTT	CCG	CCA	TGAG	AATC	AC A	CAGA	GTCT	т ст	GTAG	GG	1210
	Ala	Ser	Pro	Leu	Val	Leu	Pro	Pro									
	375					380											
	GTAT	GGTG	CG C	CGCA	TGAC	A TG	GGAG	GCGA	TGG	GGAC	GAT	GGAC	AGAG.	AC A	GAGC	GTGCA	1270
35																TGACT	1330
																ACTGC	1390
																ATGCC	1450
	TCTC																1502

	(2) INFORMATION	TOR BEQ ID NO: 56:	
	(i) SEQUE	ENCE CHARACTERISTICS:	
	(A)	LENGTH: 1349	
5	(B)	TYPE: Nucleic acid	
	(C)	STRANDEDNESS: Double	
	(D)	TOPOLOGY: Linear	
	(ii) SEQU	JENCE KIND: cDNA to mRNA	
10	(vi) ORIG	INAL SOURCE:	
	(A)	ORGANISM: Homo sapiens	
	(B)	CELL KIND: Liver	
	(D)	CLONE NAME: HP01299	
15			
	(ix) SEQU	ENCE CHARACTERISTICS:	
	(A)	CHARACTERIZATION CODE: CDS	
	(B)	EXISTENCE POSITION: 111 1064	
	(C)	CHARACTERIZATION METHOD: E	
20			
	(xi) SEQU	ENCE DESCRIPTION: SEQ ID NO: 38:	
		GAGGAA GCCGACTGCT GCCTGGTCTG CAAAGAAGTC CTTTCAAGTC	60
	TCTAGGACTG GACT	CTTCCT AAGCAAGTCC GAGAAGGAAG CACCCTCACT ATG TGG	116
25		Met Trp	
		1	
		GCC TTC GTG GGC CTG TAC TAC CTT CTG CAC TGG TAC	
	164		
		Ala Phe Val Gly Leu Tyr Tyr Leu Leu His Trp Tyr	
30	5	10 15	
		GTG GTG AGC CAC CTC CAA GAC AAG TAT GTC TTT ATC	212
		Val Val Ser His Leu Gln Asp Lys Tyr Val Phe Ile	
	20	25 30	
) E		TCG GGC TTT GGG AAC CTG CTG GCC AGA CAG CTG GAT	260
35		Ser Gly Phe Gly Asn Leu Leu Ala Arg Gln Leu Asp	
	35 CCA CCA CCC TITE	40 45 50	200
		AGA GTG CTG GCT GCG TGT CTG ACG GAG AAG GGG GCC	308
	wis will give ren	Arg Val Leu Ala Ala Cys Leu Thr Glu Lys Gly Ala	

				55	5				60	)				6.5	5	
GAG	CAC	CTG	AGG	GGC	CAG	ACC	TCI	GAC	AGG	CTO	GAC	ACG	GTO	ACC	CTG	356
G1u	Glr	ı Leu	ı Arg	Gly	Gln	Thr	Ser	Asp	Arg	, Lei	ı Glu	ı Thr	Va]	. Thi	Leu	
			70	•				75	5				80	)		
GAT	GTI	ACC	AAG	ATG	GAG	AGC	ATC	GCI	GCA	GCI	' ACI	CAG	TGG	GTG	AAG	404
Asp	Val	Thr	Lys	Met	Glu	Ser	Ile	Ala	Ala	Ala	Thr	Gln	Trp	Val	Lys	
		· 85	,				90	١				95				
GAG	CAI	GTG	GGG	GAC	AGA	GGA	CTC	TGG	GGA	CTG	GTG	AAC	AAT	GCA	GGC	452
Glu	His	Val	Gly	Asp	Arg	G1y	Leu	Trp	Gly	Leu	Val	Asn	Asn	Ala	Gly	
	100	)				105	٠.				110	1				
ATT	CTT	ACA	CCA	ATT	ACC	TTA	TGT	GAG	TGG	CTG	AAC	ACT	GAG	GAC	TCT	500
Ile	Leu	Thr	Pro	Ile	Thr	Leu	Cys	Glu	Trp	Leu	Asn	Thr	Glu	Asp	Ser	
115					120					125					130	
ATG	AAT	ATG	CTC	AAA	GTG	AAC	CTC	ATT	GGT	GTG	ATC	CAG	GTG	ACC	TTG	548
Met	Asn	Met	Leu	Lys	Val	Asn	Leu	Ile	Gly	Val	Ile	Gln	Val	Thr	Leu	
				135			•		140					145		
																596
Ser	Met	Leu	Pro	Leu	Val	Arg	Arg	Ala	Arg	Gly	Arg	Ile	Val	Asn	Val	
			150					155					160			
																644
Ser	Ser		Leu	Gly	Arg	Val	Ala	Phe	Phe	Va1	Gly	Gly	Tyr	Cys	Val	
							170					175				
																692
Ser		Tyr	Gly	Val	Glu		Phe	Ser	Asp	Ile	Leu	Arg	Arg	Glu	Ile	
	-										190					
																740
	nıs	rne	Gly	Val		Ile	Ser	Ile	Val		Pro	Gly	Tyr	Phe		
	004	4 ma														
																788
int	GIY	met			Met	Thr	GIn	Ser		Glu	Arg	Met	Lys		Ser	
TCC		C A A			4.40											
																836
пр	ьys	GIU		Pro	Lys	Hls	TIE		Glu	Thr	Tyr			Gln	Tyr	
ጥጥጥ	САТ	GCC		ጥልሮ	A A TT	A TI C	4 TD C		044	000	000			<b></b>		
																884
	p		Leu	- y I	voli	116		пÄд	GIU	ота	ren		ASN	cys	ser	
ACA	AAC		AAC	ርፓር	GTC	ልርጥ		ፐርር	A TC	CAA	<b>ር</b> ልጥ		ርሞር	<b>A</b> C <b>A</b>	<b>ጥ</b> ርር	032
	GAT Asp GAG Glu ATT Ile 115 ATG Met TCC Ser TCC Ser TCC TTC TCC TTC TTT TTT TTT The	GAT GTT Asp Val  GAG CAT Glu His 100 ATT CTT Ile Leu 115 ATG AAT Met Asn  AGC ATG Ser Met  TCC AGC Ser Ser  TCC AAG Ser Lys 180 CAA CAT Gln His 195 ACG GGA Thr Gly  TGG AAA Trp Lys  TTT GAT Phe Asp	GAT GTT ACC Asp Val Thr 85 GAG CAT GTG Glu His Val 100 ATT CTT ACA Ile Leu Thr 115 ATG AAT ATG Met Asn Met  AGC ATG CTT Ser Met Leu  TCC AGC ATT Ser Ser Ile 165 TCC AAG TAT Ser Lys Tyr 180 CAA CAT TTT Gln His Phe 195 ACG GGA ATG Thr Gly Met  TGG AAA GAA Trp Lys Glu  TTT GAT GCC Phe Asp Ala 245	Glu Gln Leu Arg 70 GAT GTT ACC AAG Asp Val Thr Lys 85 GAG CAT GTG GGG Glu His Val Gly 100 ATT CTT ACA CCA Ile Leu Thr Pro 115 ATG AAT ATG CTC Met Asn Met Leu AGC ATG CTT CCT Ser Met Leu Pro 150 TCC AGC ATT CTG Ser Ser Ile Leu 165 TCC AAG TAT GGA Ser Lys Tyr Gly 180 CAA CAT TTT GGG Gln His Phe Gly 195 ACG GGA ATG ACA Thr Gly Met Thr TGG AAA GAA GCC Trp Lys Glu Ala 230 TTT GAT GCC CTT Phe Asp Ala Leu 245	GAG CAG CTG AGG GGG Glu Gln Leu Arg Gly 70  GAT GTT ACC AAG ATG Asp Val Thr Lys Met 85  GAG CAT GTG GGG GAC Glu His Val Gly Asp 100  ATT CTT ACA CCA ATT Ile Leu Thr Pro Ile 115  ATG AAT ATG CTC AAA Met Asn Met Leu Lys 135  AGC ATG CTT CCT TTG Ser Met Leu Pro Leu 150  TCC AGC ATT CTG GGA Ser Ser Ile Leu Gly 165  TCC AAG TAT GGA GTG Ser Lys Tyr Gly Val 180  CAA CAT TTT GGG GTG Gln His Phe Gly Val 195  ACG GGA ATG ACA AAC Thr Gly Met Thr Asn 215  TGG AAA GAA GCC CCC Trp Lys Glu Ala Pro 230  TTT GAT GCC CTT TAC Phe Asp Ala Leu Tyr 245	Glu Gln Leu Arg Gly Gln 70  GAT GTT ACC AAG ATG GAG Asp Val Thr Lys Met Glu 85  GAG CAT GTG GGG GAC AGA Glu His Val Gly Asp Arg 100  ATT CTT ACA CCA ATT ACC Ile Leu Thr Pro Ile Thr 115 120  ATG AAT ATG CTC AAA GTG Met Asn Met Leu Lys Val 135  AGC ATG CTT CCT TTG GTG Ser Met Leu Pro Leu Val 150  TCC AGC ATT CTG GGA AGA Ser Ser Ile Leu Gly Arg 165  TCC AAG TAT GGA GTG GAA Ser Lys Tyr Gly Val Glu 180  CAA CAT TTT GGG GTG AAA Gln His Phe Gly Val Lys 195 200  ACG GGA ATG ACA AAC ATG Thr Gly Met Thr Asn Met 215  TGG AAA GAA GCC CCC AAG TTP Lys Glu Ala Pro Lys 230  TTT GAT GCC CTT TAC AAT Phe Asp Ala Leu Tyr Asn 245	GAG CAG CTG AGG GGC CAG ACC Glu Gln Leu Arg Gly Gln Thr 70  GAT GTT ACC AAG ATG GAG AGG Asp Val Thr Lys Met Glu Ser 85  GAG CAT GTG GGG GAC AGA GGA Glu His Val Gly Asp Arg Gly 100 105  ATT CTT ACA CCA ATT ACC TTA Ile Leu Thr Pro Ile Thr Leu 115 120  ATG AAT ATG CTC AAA GTG AAC Met Asn Met Leu Lys Val Asn 135  AGC ATG CTT CCT TTG GTG AGG Ser Met Leu Pro Leu Val Arg 150  TCC AGC ATT CTG GGA AGA GTT Ser Ser Ile Leu Gly Arg Val 165  TCC AAG TAT GGA GTG GAA GCC Ser Lys Tyr Gly Val Glu Ala 180 185  CAA CAT TTT GGG GTG AAA ATC Gln His Phe Gly Val Lys Ile 195 200  ACG GGA ATG ACA AAC ATG ACA Thr Gly Met Thr Asn Met Thr 215  TGG AAA GAA GCC CCC AAG CAT Trp Lys Glu Ala Pro Lys His 230  TTT GAT GCC CTT TAC AAT ATC Phe Asp Ala Leu Tyr Asn Ile	GAG CAG CTG AGG GGC CAG ACG TCT Glu Gln Leu Arg Gly Gln Thr Ser 70  GAT GTT ACC AAG ATG GAG AGC ATG ASP VA1 Thr Lys Met Glu Ser Ilee 85 90  GAG CAT GTG GGG GAC AGA GGA CTG Glu His Va1 Gly Asp Arg Gly Leu 100 105  ATT CTT ACA CCA ATT ACC TTA TGT Ile Leu Thr Pro Ile Thr Leu Cys 115 120  ATG AAT ATG CTC AAA GTG AAC CTG Met Asn Met Leu Lys Va1 Asn Leu 135  AGC ATG CTT CCT TTG GTG AGG AGA Ser Met Leu Pro Leu Va1 Arg Arg 150  TCC AGC ATT CTG GGA AGA GTT GCT Ser Ser Ile Leu Gly Arg Va1 Ala 165 170  TCC AAG TAT GGA GTG GAA GCC TTT Ser Lys Tyr Gly Va1 Glu Ala Phe 180 185  CAA CAT TTT GGG GTG AAA ATC AGC GIn His Phe Gly Va1 Lys Ile Ser 195 200  ACG GGA ATG ACA AAC ATG ACA CAG TTT GTG AAA GAC CAG TTT GLY Met Thr Asn Met Thr Gln 215  TGG AAA GAA GCC CCC AAG CAT ATT TTP Lys Glu Ala Pro Lys His Ile 230  TTT GAT GCC CTT TAC AAT ATC ATG Phe Asp Ala Leu Tyr Asn Ile Met 245 250	GAG CAG CTG AGG GGC CAG ACG TCT GAG Clu Gln Leu Arg Gly Gln Thr Ser Asp 70 72  GAT GTT ACC AAG ATG GAG AGC ATC GCT Asp Val Thr Lys Met Glu Ser Ile Ala 85 90  GAG CAT GTG GGG GAC AGA GGA CTC TGG Glu His Val Gly Asp Arg Gly Leu Trp 100 105  ATT CTT ACA CCA ATT ACC TTA TGT GAG Ile Leu Thr Pro Ile Thr Leu Cys Glu 115 120  ATG AAT ATG CTC AAA GTG AAC CTC ATT Met Asn Met Leu Lys Val Asn Leu Ile 135  AGC ATG CTT CCT TTG GTG AGG AGA GCA Ser Met Leu Pro Leu Val Arg Arg Ala 150 155  TCC AGC ATT CTG GGA AGA GTG GCC TTC Ser Ser Ile Leu Gly Arg Val Ala Phe 165 170  TCC AAG TAT GGA GTG GAA GCC TTT TCA Ser Lys Tyr Gly Val Glu Ala Phe 180 185  CAA CAT TTT GGG GTG AAA ATC ACC ATA Gln His Phe Gly Val Lys Ile Ser Ile 195 200  ACG GGA ATG ACA ACC AAC ATG ACA CAG TCC Thr Gly Met Thr Asn Met Thr Gln Ser 215  TGG AAA GAA GAC CCC CCC AAG CAT ATT AAG Trp Lys Glu Ala Pro Lys His Ile Lys 230 235  TTT GAT GCC CTT TAC AAT ATC ATG AAG Phe Asp Ala Leu Tyr Asn Ile Met Lys 245 250	GAG CAG CTG AGG GGC CAG ACG TCT GAC AGG Glu Gln Leu Arg Gly Gln Thr Ser Asp Arg 70 75  GAT GTT ACC AAG ATG GAG AGC ATC GCT GCA Asp Val Thr Lys Met Glu Ser Ile Ala Ala 85 90  GAG CAT GTG GGG GAC AGA GGA CTC TGG GGA Glu His Val Gly Asp Arg Gly Leu Trp Gly 100 105  ATT CTT ACA CCA ATT ACC TTA TGT GAG TGG Ile Leu Thr Pro Ile Thr Leu Cys Glu Trp 115 120  ATG AAT ATG CTC AAA GTG AAC CTC ATT GGT Met Asn Met Leu Lys Val Asn Leu Ile Gly 135 140  AGC ATG CTT CCT TTG GTG AGG AGA GCA CGG Ser Met Leu Pro Leu Val Arg Arg Ala Arg 150 155  TCC AGC ATT CTG GGA AGA GTT GCT TTC TTT Ser Ser Ile Leu Gly Arg Val Ala Phe Phe 165 170  TCC AAG TAT GGA GTG GAA GCC TTT TCA GAT Ser Lys Tyr Gly Val Glu Ala Phe Ser Asp 180 185  CAA CAT TTT GGG GTG AAA ATC AGC ATA GTT Gln His Phe Gly Val Lys Ile Ser Ile Val 195 200  ACG GGA ATG ACA ACA ACC ATG ACA CAG TCC TTA Thr Gly Met Thr Asn Met Thr Gln Ser Leu 215 220  TGG AAA GAA GCC CCC AAG CAT ATT AAG GAG TTP Lys Glu Ala Pro Lys His Ile Lys Glu 230 235  TTT GAT GCC CTT TAC AAT ATC ATG AAG GAA Phe Asp Ala Leu Tyr Asn Ile Met Lys Glu 245 250	GAG CAG CTG AGG GGC CAG ACG TCT GAC AGG CTG Glu Gln Leu Arg Gly Gln Thr Ser Asp Arg Leu 70 75  GAT GTT ACC AAG ATG GAG AGC AGC GCT GCA GAG Asp Val Thr Lys Met Glu Ser Ile Ala Ala Ala 85 90  GAG CAT GTG GGG GAC AGA GGA CTC TGG GGA CTG Glu His Val Gly Asp Arg Gly Leu Trp Gly Leu 100 105  ATT CTT ACA CCA ATT ACC TTA TGT GAG TGG CTG Ile Leu Thr Pro Ile Thr Leu Cys Glu Trp Leu 115 120 125  ATG AAT ATG CTC AAA GTG AAC CTC ATT GGT GGG Met Asn Met Leu Lys Val Asn Leu Ile Gly Val 135 140  AGC ATG CTT CCT TTG GTG AGG AGA GCA CGG GGA Ser Met Leu Pro Leu Val Arg Arg Ala Arg Gly 150  TCC AGC ATT CTG GGA AGA GTT GCT TTC TTT GTA Ser Ser Ile Leu Gly Arg Val Ala Phe Phe Val 165 170  TCC AAG TAT GGA GTG GAA GCC TTT TCA GAT ATT Ser Lys Tyr Gly Val Glu Ala Phe Ser Asp Ile 180 185  CAA CAT TTT GGG GTG AAA ATC AGC ATA GTT GAA Gln His Phe Gly Val Lys Ile Ser Ile Val Glu 195 200 205  ACG GGA ATG ACA ACA ACA CTG ACA CAG TCC TTA GAG Thr Gly Met Thr Asn Met Thr Gln Ser Leu Glu 215 200  ACG GAAA GAA GCC CTC AAG ATC ACA CAG GAA GCC TTP Lys Glu Ala Pro Lys His Ile Lys Glu Thr 230 235  TTT GAT GCC CTT TAC AAT ATC ATG AAG GAA GGG Phe Asp Ala Leu Tyr Asn Ile Met Lys Glu Gly 245 250	GAG CAG CTG AGG GGC CAG ACG TCT GAC AGG CTG GAC Glu Gln Leu Arg Gly Gln Thr Ser Asp Arg Leu Glu 70 75  GAT GTT ACC AAG ATG GAG AGC ATC GCT GCA GCT ACT Asp Val Thr Lys Met Glu Ser Ile Ala Ala Ala Thr 85 90  GAG CAT GTG GGG GAC AGA GGA CTC TGG GGA CTG GTG Glu His Val Gly Asp Arg Gly Leu Trp Gly Leu Val 100 105 110  ATT CTT ACA CCA ATT ACC TTA TGT GAG TGG CTG AAC Ile Leu Thr Pro Ile Thr Leu Cys Glu Trp Leu Asn 115 120 125  ATG AAT ATG CTC AAA GTG AAC CTC ATT GGT GTG ATC Met Asn Met Leu Lys Val Asn Leu Ile Gly Val Ile 135 140  AGC ATG CTT CCT TTG GTG AGG AGA GCA CGG GGA AGA Ser Met Leu Pro Leu Val Arg Arg Ala Arg Gly Arg 150 155  TCC AGC ATT CTG GGA AGA GTT GCT TTC TTT GTA GGA Ser Ser Ile Leu Gly Arg Val Ala Phe Phe Val Gly 165 170  TCC AAG TAT GGA GTG GAA GCC TTT TCA GAT ATC Ser Lys Tyr Gly Val Glu Ala Phe Ser Asp Ile Leu 180 185 190  CAA CAT TTT GGG GTG AAA ATC AGC ATA GTT GAT CTG Gln His Phe Gly Val Lys Ile Ser Ile Val Glu Pro 195 200 205  ACG GGA AAA GAA GCC CCC AAG CAT ATT AAC GAA ACC TTP Lys Glu Ala Pro Lys His Ile Lys Glu Thr Tyr 230 235  TTT GAT GCC CTT TAC AAT ATC ATG AAG GAA GGC CTG Phe Asp Ala Leu Tyr Asn Ile Met Lys Glu Gly Leu 245 250	GAG CAG CTG AGG GGC CAG ACG TCT GAC AGG CTG GAG ACG Glu Gln Leu Arg Gly Gln Thr Ser Asp Arg Leu Glu Thr  70	GAG CAG CTG AGG GGC CAG ACG TCT GAC AGG CTG GAG ACG CTG GLU GLU GLU LEU ARG GLU GLU THY VAIL AGG GTG GTG GAG ACG GTG GAT GTG AGG GTT ACC AAG ATG GAG AGG ATG GCT GCT GCA GCT ACT CAG TGG ASP VAI THY LYS MET GLU SET ILE ALA ALA ALA THY GLU THY 8.5 PO 90 95 PO 9	GAG CAG CTG AGG GGC CAG ACG TCT GAC AGG CTG GAG ACG GTG ACG GLU Glu Glu Leu Arg Gly Glu Thr Ser Asp Arg Leu Glu Thr Val Thr 70	GAG CAG CTG AGG GGC CAG ACG TCT CAC AGG CTG GAG ACG GTG ACC CTC Glu Gln Leu Arg Gly Gln Thr Ser Asp Arg Leu Glu Thr Val Thr Leu 70 75 80 80 76 AGG GTG AGG AGG GTG AAG AAG ATG GAT ACC CAG GTG GAT ACC CAG TGG GAG AAG AAF CAG AAG AAF CAG AGG AGG AGG AGG AGG AGG AGG AGG AGG

124

	Thr Asn	Leu	Asn	Leu	Val	Thr	Asp	Cys	Met	Glu	His	Ala	Leu	Thr	Ser	
	260	)				265					270					
	GTG CAT	CCG	CGA	ACT	ĊGA	TAT	TCA	GCT	GGC	TGG	GAT	GCT	AAA	TTT	TTC	980
	Val His	Pro	Arg	Thr	Arg	Tyr	Ser	Ala	Gly	Trp	Asp	Ala	Lys	Phe	Phe	
5	275				280					285					290	
	TTC ATC															1028
	Phe Ile	Pro	Leu		Tyr	Leu	Pro	Thr		Leu	Ala	Asp	Tyr		Leu	
				295					300					305		
1.0	ACT AGA										TAAA	\GAA/	AAC 1	rggg:	rtggt	1080
10	Thr Arg	Ser		Pro	Lys	Pro	Ala		Ala	Val						
	CCMMCMM	004	310					315								
	GCTTCTT															
	ACCCAGG AGTACTA															
15	ACTATTT															
	TATTAAA							. 116	TOIR	MAG	IGAR	IICAI	.11 6	,110,	16001	1320 1349
							••••									1349
	(2) INF	ORMAT	NOI	FOR	SEQ	ID N	10: 3	9:								
20	(:	i) SE	EQUEN	CE C	HARA	CTER	ISTI	CS:								
			(A)	LENG	TH:	1643	3									
			(B)	TYPE	: Nu	clei	.c ac	id								
			(C)	STRA	NDED	NESS	: Do	uble								
			(D)	TOPO	LOGY	: Li	near									
25	(:	Li) S	EQUE	NCE	KIND	: cD	NA t	o mR	NA							
	(1	7i) 0														
				ORGA				•	ns							
2.0				CELL												
30			(D)	CLON	E NA	ME:	HP01	347								
	( ;	lx) S	FOIIF	NCF :	CHAD	ልሮጥፑ	DTCT	TCC.								
	(-			CHAR.					F. C	ne						
				EXIS												
35			,							713						
J -			(C)	CHAR	ACTE	RIZA	TTON	MET	• пон	F.						

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 39:

	AAC	ATCI	GGG	GACA	.GCGG	GA A	AAC	ATG	AGT	GAÇ	TCC	AAG	GAA	CCA	AGG	GTG	51
								Met	Ser	Asp	Ser	Lys	Glu	Pro	Arg	Val	•
		٠						1				5					
	CAG	CAG	CTG	GGC	CTC	CTG	GGG	TGT	CTI	GGC	CAT	GGC	GCC	CTG	GTO	CTG	99
5	Gln	Gln	Leu	Gly	Leu	Leu	Gly	Cys	Leu	Gly	His	Gly	Ala	Let	ı Val	Leu	
	10					15					20	)				25	
	CAA	CTC	CTC	TCC	TTC	ATG	CTC	TTG	GCT	GGG	GTC	CTG	GTG	GCC	ATC	CTT	147
	Gln	Leu	Leu	Ser	Phe	Met	Leu	Leu	Ala	Gly	Val	Leu	Val	Ala	Ile	Leu	
		•			30					35					40	)	
10	GTC	CAA	GTG	TCC	AAG	GTC	ccc	AGC	TCC	CTA	AGT	CAG	GAA	CAA	TCC	GAG	195
	Val	Gln	Val	Ser	Lys	Val	Pro	Ser	Ser	Leu	Ser	Gln	G1u	Gln	Ser	Gļu	
				45				•	50					55			
	CAA	GAC	GCA	ATC	TAC	CAG	AAC	CTG	ACC	CAG	CTT	AAA	GCT	GCA	GTG	GGT	243
	Gln	Asp	Ala	Ile	Tyr	Gln	Asn	Leu	Thr	Gln	Leu	Lys	Ala	Ala	Val	Gly	
15			60					65					70				
	GAG	CTC	TCA	GAG	AAA	TCC	AAG	CTG	CAG	GAG	ATC	TAC	CAG	GAG	CTG	ACC	291
	G1u	Leu	Ser	Glu	Lys	Ser	Lys	Leu	Gln	Glu	Ile	Tyr	Gln	Glu	Leu	Thr	
		75					80					85					
	CAG	CTG	AAG	GCT	GCA	GTG	GGT	GAG	TTG	CCA	GAG	AAA	TCC	AAG	CTG	CAG	339
20	Gln	Leu	Lys	Ala	Ala	Val	Gly	Glu	Leu	Pro	Glu	Lys	Ser	Lys	Leu	Gln	
	90					95					100					105	
	GAG	ATC	TAC	CAG	GAG	CTG	ACC	CGG	CTG	AAG	GCT	GCA	GTG	GGT	GAG	TTG	387
	Glu	Ile	Tyr	Gln	Glu	Leu	Thr	Arg	Leu	Lys	Ala	-Ala	Val	Gly	Glu	Leu	
					110					115					120		
25	CCA	GAG	AAA	TCC	AAG	CTG	CAG	GAG	ATC	TAC	CAG	GAG	CTG	ACC	CGG	CTG	435
	Pro	Glu	Lys	Ser	Lys	Leu	Gln	Glu	Ile	Tyr	Gln	Glu	Leu	Thr	Arg	Leu	
				125					130					135			
						GAG											483
	Lys	Ala		Val	Gly	Glu	Leu	Pro	Glu	Lys	Ser	Lys	Leu	Gln	Glu	Ile	
30			140					145					150				
						CGG											531
	Tyr	Gln	Glu	Leu	Thr	Arg	Leu	Lys	Ala	Ala	Val	G1y	Glu	Leu	Pro	Glu	
		155					160					165					
						GAG											579
35		Ser	Lys	Leu	Gln	Glu	Ile	Tyr	Gln	Glu		Thr	Glu	Leu	Lys		
	170					175					180					185	
•	_					CCA											627
	Ala	Val	Glv	Glu	Leu	Pro	Glu	Lvs	Ser	Lvs	Leu	Gln	Glu	Tle	Tvr	Gln	

					190					195					200		
	GAG	CTG	ACC	CAG	CTG	AAG	GCT	GCA	GTG	GGT	GAG	TTG	CCA	GAC	CAG	TCC	67
	Glu	Leu	Thr	Gln	Leu	Lys	Ala	Ala	Val	Gly	Glu	Leu	Pro	Asp	Gln	Ser	
				205					210					215			
5	AAG	CAG	CAG	CAA	ATC	TAT	CAA	GAA	CTG	ACC	GAT	TTG	AAG	ACT	GĊA	TTT	723
	Lys	Gln	Gln	Gln	Ile	Tyr	Gln	Glu	Leu	Thr	Asp	Leu	Lys	Thr	Ala	Phe	
			220					225					230				
	GAA	CGC	CTG	TGC	CGC	CAC	TGT	CCC	AAG	GAC	TGG	ACA	TTC	TTC	CAA	GGA	771
	Glu	Arg	Leu	Cys	Arg	His	Cys	Pro	Lys	Asp	Trp	Thr	Phe	Phe	Gln	Gly	
10		2,35					240					245					
	AAC	TGT	TAC	TTC	ATG	TCT	AAC	TCC	CAG	CGG	AAC	TGG	CAC	GAC	TCC	GTC	819
	Asn	Cys	Tyr	Phe	Met	Ser	Asn	Ser	Gln	Arg	Asn	Trp	His	Asp	Ser	Val	
	250					255					260					265	
	ACC	GCC	TGC	CAG	GAA	GTG	AGG	GCC	CAG	CTC	GTC	GTA	ATC	AAA	ACT	GCT	867
15	Thr	Ala	Cys	Gln	Glu	Val	Arg	Ala	Gln	Leu	Val	Val	Ile	Lys	Thr	Ala	
					270					275					280		
	GAG	GAG	CAG	CTT	CCA	GCG	GTA	CTG	GAA	CAG	TGG	AGA	ACC	CAA	CAA		912
	Glu	Glu	Gln	Leu	Pro	Ala	Val	Leu	Glu	Gln	Trp	Arg	Thr	G1n	Gln		
				285					290			_		295			
20	TAGO	CGGGA	AT G	SAAGA	CTGT	G CG	GAAT	TTAG	TGG	CAGI	GGC	TGGA	ACGA	CA A	TCGA	TGT	970
	GAC	TTGA	CA A	TTAC	TGGA	T CT	GCAA	AAAG	ccc	GCAG	CCT	GCTT	'CAGA	GA C	GAAT	AGTTG	1030
	TTTC	CCTG	CT A	GCCT	CAGO	C TC	CATT	GTGG	TAT	'AGCA	GAA	CTTC	ACCC	AC I	TGTA	AGCCA	1090
	GCGC	TTCT	TC 1	CTCC	ATCC	T TG	GACC	TTCA	CAA	ATGC	CCT	GAGA	CGGT	TC T	CTGT	TCGAT	1150
	TTTT	CATO	cc c	TATG	AACC	T GG	GTCT	TATT	CTG	TCCT	TCT	GATG	CCTC	CA A	GTTT.	CCCTG	1210
25	GTGT	'AGAG	CT I	GTGT	TCTT	G GC	CCAT	CCTT	GGA	GCTT	TAT	AAGT	GACC	TG A	.GTGG	GATGC	1270
	ATTI	'AGGG	GG C	GGGC	TTGG	T AT	GTTG	TATG	AAT	CCAC	TCT	CTGT	TCCT	тт т	GGAG	ATTAG	1330
	ACTA	TTTG	GA I	'TCAT	GTGT	A GC	TGCC	CTGT	CCC	CTGG	GGC	TTTA	TCTC	AT C	CATG	CAAAC	1390
																CTGGT	1450
																CCAAT	1510
30												,				CTTTG	1570
																GGTAA	1630
		TGTA															1643

## 35 (2) INFORMATION FOR SEQ ID NO: 40:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 729
  - (B) TYPE: Nucleic acid

			(0)	, 511	CUIDI	ביווענ	.J. I	JOUDI	. e							
			(D)	TOF	POLOG	Y: L	ines	ır								
		(ii)	SEQU	JENCE	KIN	ID: c	DNA	to m	ıRNA							
5		(vi)	ORIG	INAL	SOU	RCE:										
			(A)	ORG	ANIS	M: <i>H</i>	ото	sapi	ens							
			(B)	CEL	L KI	ND:	Ston	ach	cano	er						
			(D)	CLO	NE N	IAME:	HPO	1440	)							
10		(ix)	SEQU	ENCE	СНА	RACT	ERIS	TICS	:							
			(A)	CHA	RACT	ERIZ	ATIO	N CO	DE:	CDS						
		* -	(B)	EXI	STEN	CE P	OSIT	ION:	38.	. 63	1					
			(C)	CHA	RACT	ERIZ	OITA	N ME	THOD	: E						
15	(	(xi)	SEQU	ENCE	DES	CRIP	TION	: SE	Q ID	NO:	40:					
	A COMMON	000			<b>.</b>											_
	ACTTTCA	ICTC	ACCG	CCTG	TC C	TTCC	TGAC	A CC	TCAC							5:
											•	s Tn	r GI	-	s Cys	
20	GCC CGC	: ጥርጥ	ርጥር	GGG	ርሞር	ሞርር	CTC	ለ ጥጥ	۸۵۵		l TCC	ርሞር	CTC		5 ልጥጥ	10:
20	Ala Arg															
			10	01)	204	501		15	1111	Бец	oy s		20	0,3	110	
	GTG GCC			СТС	CTG	CTG			ААТ	GGG	GAG			TGG	ACC	151
	Val Ala															,
25		25					30			,		35				
	AAC ACC	AAC	CAT	CTC	AGC	TTG	CAA	GTC	TGG	CTC	ATG	GGC	GGC	TTC	ATT	199
	Asn Thr	Asn	His	Leu	Ser	Leu	Gln	Val	Trp	Leu	Met	Gly	Gly	Phe	Ile	
	40	)				45					50					
	GGC GGG	GGC	CTA	ATG	GTA	CTG	TGT	CCG	GGG	ATT	GCA	GCC	GTT	CGG	GCA	247
30	Gly Gly	Gly	Leu	Met	Val	Leu	Cys	Pro	Gly	Ile	Ala	Ala	Val	Arg	Ala	
	55				60					65					70	
	GGG GGC	AAG	GGC	TGC	TGT	GGT	GCT	GGG	TGC	TGT	GGA	AAC	CGC	TGC	AGG	295
	Gly Gly	Lys	Gly	Cys	Cys	Gly	Ala	Gly	Cys	Cys	Gly	Asn	Arg	Cys	Arg	
				75	•				80					85		
35	ATG CTG															343
	Met Leu	Arg		Val	Phe	Ser	Ser	Ala	Phe	Gly	Val	Leu	Gly	Ala	Ile	
			90					95					100			
	TAC TGC	CTC	TCG	GTG	TCT	GGA	CCT	CCC	CTC	CCA	ጥ ል ል	CCA	CCC	ACA	TGC	301

	Tyr	Cys	Leu	Ser	Val	Ser	Gly	Ala	Gly	Leu	Arg	Asn	Gly	Pro	Arg	Cys	
			105					110					115				
	TTA	ATG	AAC	GGC	GAG	TGG	GGC	TAC	CAC	TTC	GAA	GAC	ACC	GCG	GGA	GCT	439
	Leu	Met	Asn	Gly	Glu	Trp	Gly	Tyr	His	Phe	Glu	Asp	Thr	Ala	Gly	Ala	
5		120					125					130					
															CCT		487
	Tyr	Leu	Leu	Asn	Arg	Thr	Leu	Trp	Asp	Arg	Сув	Glu	Ala	Pro	Pro	Arg	
	135					140					145					150	
															GCC		535
10	Val	Val	Pro	Trp		Val	Thr	Leu	Phe	Ser	Leu	Leu	Val	Ala	Ala	Ser	
					155					160					165		
															ACC		583
	Cys	Leu	Glu		Val	Leu	Cys	Gly	Ile	Gln	Leu	Val	Asn	Ala	Thr	Ile	
				170					175					180			
15															CAC	TG	630
	Gly	Val		Cys	Gly	Asp	Cys		Lys	Lys	Gln	Asp		Pro	His		
	1000		185					190					195				
												ACGC	CTAC	CT G	GCTC	GCTCA	690
20	CICC	CTTG	our c	GGT	IGAAT	'A AA	CTGC	TTTG	GGC	TCTC	TT						729
				,													
	(2)	INFO	RMAT	ON	FOR	SEQ	ID N	10: 4	1:								
	\- <i>\</i>	_				HARA											
				•		TH:											
25	٠.			(B)	TYPE	: Nu	clei	c ac	id								
				(C)	STRA	NDED	NESS	: Do	uble	!							
				(D)	TOPO	LOGY	: Li	.near	•								
		(i	i) S	EQUE	NCE	KIND	: cD	NA t	o mR	.NA							
30		(v	i) 0	RIGI	NAL	SOUR	CE:										
						NISM											
						KIN				ance	r						
				(D)	CLON	E NA	ME:	HP01	526								
35		(i	x) S	EQUE	NCE	CHAR	ACTE	RIST	ICS:								
				(A)	CHAR	ACTE	RIZA	TION	COD	E: C	DS						
				(B)	EXIS	TENC	E PO	SITI	ON:	84	749						
				/C\	CHAD	ለ <b>ር</b> ጥ ፱	D T 7 A	MT ON	MEM	110D	-						

## (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 41:

	GAG	CCGC.	AGG	TCTG	GGCT	GC A	GTAG	GTCC	C GG	CAA	CCGC	A GG	CTCG	CGGC	GGG	CGC	TGGG	60
	CGC	GGGA	TCC	GACT	CTAG	TC G	TA A	TG G	AG G	CG (	GC (	GGC	TTT	CTG	GAC	TCG	CTC	113
5							М	et G	lu A	la (	3ly (	Gly	Phe	Leu .	Asp	Ser	Leu	
								1				. 5					10	
	ATT	TAC	GGA	GCA	TGC	GTG	GTC	TTC	ACC	CTT	r GG	CAT	G TT	C TC	C GC	C G	3C	161
	Ile	Tyr	Gly	Ala	Ċys	Val	Val	Phe	Thr	Let	1 G1	y Me	t Ph	e Se	r Ala	a G	ly	
					15					20	)				2	5		
10	CTC	TCG	GAC	CTC	AGG	CAC	ATG	CGA	ATG	ACC	CG	G AG	r GT	G GA	C AA	G G:	rc	209
	Leu	Ser	Asp	Leu	Arg	His	Met	Arg	Met	Thi	: Arg	g Se	r Vai	l Ası	p Ası	a Ve	<b>a</b> l	
				30					35					4 (	0			
	CAG	TTC	CTG	CCC	TTT	CTC	ACC	ACG	GAA	GTC	AA(	AA C	CTC	G GG(	C TG	G C	ľG	257
	G1n	Phe	Leu	Pro	Phe	Leu	Thr	Thr	Glu	Val	. Ası	n Ası	n Lei	ı G13	y Tr	Le	eu	
15			45					50					5	5				
	AGT	TAT	GGG	GCT	TTG	AAG	GGA	GAC	GGG	ATC	CTC	CATO	GTO	GTO	C AAC	CAC	CA	305
	Ser	Tyr	Gly	Ala	Leu	Lys	Gly	Asp	Gly	Ile	Leu	ı Ile	e Val	L Val	L Ası	1 Th	ır	
		60					65					70	)					
	GTG	GGT	GCT	GCG	CTT	CAG	ACC	CTG	TAT	ATC	TTO	GCA	A TAT	CTC	CAT	TA	C	353
20	Val	G1y	Ala	Ala	Leu	Gln	Thr	Leu	Tyr	Ile	Leu	ı Ala	ту т	Let	ı His	ту	r	
	75			•		80					85		•			-	0	
	TGC	CCT	CGG	AAG	CGT	GTT	GTG	CTC	CTA	CAG	ACT	GCA	ACC	CTC	CTA	GG	G	401
	Cys	Pro	Arg	Lys	Arg	Val	Val	Leu	Leu	Gln	Thr	Ala	Thi	Leu	ı Let	ı G1	. <b>y</b>	
					95					100					105			
25															) AAC			449
	Val	Leu	Leu		Gly	Tyr	Gly	Tyr		Trp	Lev	Let	ı Val		Asr	Pr	о.	
				110					115					120				
															ACC			497
20	Glu	Ala		Leu	Gln	Gln	Leu		Leu	Phe	Cys	Ser			Thr	. 11	е	
30	400	4.50	125	0.00	<b></b>			130					135				_	
							•								CAA			545
	ser		Tyr	Leu	Ser	Pro		Ala	Asp	Leu	Ala	•		. Ile	Gln	Th	r	
		140	400				145					150					_	
2 5															CTT			593
35		ser	inr	GIN	cys		ser	Tyr	Pro	Leu			ALA	rnr	Leu			
	155	ጥርጥ	ccc	TCC	TCC	160	CTC	ጥልጥ	ccc	mmm	165			C 4 m		17 • • •		61.7
															CCC Pro			641
	T 11T	SET	UTG	OCT	trb	∪yS	Leu	TAL	GTA	rne	AIR	ren	LATE	nsp	PLO	' ↓y		

	175 180 185	
	ATC ATG GTG TCC AAC TTT CCA GGA ATC GTC ACC AGC TTT ATC CGC TTC	689
	Ile Met Val Ser Asn Phe Pro Gly Ile Val Thr Ser Phe Ile Arg Phe	
	190 195 200	
5	TGG CTT TTC TGG AAG TAC CCC CAG GAG CAA GAC AGG AAC TAC TGG CTC	737
	Trp Leu Phe Trp Lys Tyr Pro Gln Glu Gln Asp Arg Asn Tyr Trp Leu	
	205 210 215	
	CTG CAA ACC TGAGGCTGCT CATCTGACCA CTGGGCCACCT TAGTGCCAAC CTGA	790
	Leu Gln Thr	
10	220	
	ACCAAAGAGA CCTCCTTGTT TCAGCTGGGC CTGCTGTCCA GCTTCCCAGG TGCAGTGGGT	850
	TGTGGGAACA AGAGATGACT TTGAGGATAA AAGGACCAAA GAAAAAGCTT TACTTAGATG	910
	ATTGATTGGG GCCTAGGAGA TGAAATCACT TTTTATTTTT TAGAGATTTT TTTTTTTAAT	970
	TTTGGAGGTT GGGGTGCAAT CTTTAGAATA TGCCTTAAAA GGCCGGGCGC GGTGGCTCAC	1030
15	GCCTGTAATC CCAGCACTTT GGGAGGCCAA GGTGGGCGGA TCGCCTGAGG TCAGGAGTTC	1090
	AAGACCAACC TGACTAACAT GGTGAAACCC CATCTCTACT AAAAATACAA AATTAGCCAG	1150
	GCATGATGCC ACATGCCTGT AATCCCAGAT ACTTGGGAGG CTGAGGCAGG AGAATTGCTT	1210
	GAACCCAGGA GGTGGAGGTT GCAGTGAGCT GAGATCGTGC CATTGTGATA TGAATATGCC	1270
20	TTATATGCTG ATATGAATAT GCCTTAAAAT AAAGTGTTCC CCACCCCTGC CC	1322
20		
	(2) INFORMATION FOR SEQ ID NO: 42:	
	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 3045	
25	(B) TYPE: Nucleic acid	
	(C) STRANDEDNESS: Double	
	(D) TOPOLOGY: Linear	
	(ii) SEQUENCE KIND: cDNA to mRNA	
30	(vi) ORIGINAL SOURCE:	
	(A) ORGANISM: Homo sapiens	
	(B) CELL KIND: Stomach cancer	
	(D) CLONE NAME: HP10230	
35	(ix) SEQUENCE CHARACTERISTICS:	
	(A) CHARACTERIZATION CODE: CDS	
	(B) EXISTENCE POSITION: 191 946	
	(C) CHARACTERIZATION METHOD: E	

## (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 42:

	GTT	TCGC	CTC	AGAA	GGCI	GC C	TCGC	TGGT	OO. OO	TAA	CGGT	GGC	CGCCA	CGT	CCGC	CCGTC	T 60
	CCG	CCTT	CTG	CATO	GCGG	CT I	CGGC	GGC	T CC	CACCI	'AGAC	ACC	TAAC	AGT	CGCG	GAGCC	G 120
5	GCC	GCGT	CGT	GAGG	GGGT	CG G	CACG	GGGA	G TC	CGGGC	GGTC	TTG	TGCA	TCT	TGGC	TACCT	G 180
	TGG	GTCG.	AAG	ATG	TCG	GAC	ATC	GGA	GAC	TGG	TTC	AGG	AGC	ATC	CCG	GCG	229
				Met	Ser	Asp	Ile	Gly	Asp	Trp	Phe	Arg	Ser	Ile	Pro	Ala	
				1				5					10				
	ATC	ACG	CGC	TAT	TGG	TTC	GCC	GCC	ACC	GTC	GCC	GTG	ccc	TTG	GTC	GGC	277
10	Ile	Thr	Arg	Tyr	Trp	Phe	Ala	Ala	Thr	. Val	Ala	Val	Pro	Leu	Val	Gly	
		15					20					25	i				
	AAA	CTC	GGC	CTC	ATC	AGC	CCG	GCC	TAC	CTC	TTC	CTC	TGG	ccc	GAA	GCC	325
	Lys	Leu	G1y	Leu	Ile	Ser	Pro	Ala	Tyr	Leu	Phe	Leu	Trp	Pro	Glu	Ala	
	30					35					40					45	
15	TTC	CTT	TAT	CGC	TTT	CAG	ATT	TGG	AGG	CCA	ATC	ACT	GCC	ACC	TTT	TAT	373
	Phe	Leu	Tyr	Arg	Phe	Gln	Ile	Trp	Arg	Pro	Ile	Thr	Ala	Thr	Phe	Tyr	
					50					55					60		
	TTC	CCT	GTG	GGT	CCA	GGA	ACT	GGA	TTT	CTT	TAT	TTG	GTC	AAT	TTA	TAT	421
	Phe	Pro	Val	Gly	Pro	Gly	Thr	G1y	Phe	Leu	Tyr	Leu	Val	Asn	Leu	Tyr	
20	•			65					70	)				75			
														TTT			469
	Phe	Leu	Tyr	Gln	Tyr	Ser	Thr	Arg	Leu	Glu	Thr	Gly	Ala	Phe	Asp	G1y	
			80					85					90				
														ATT			517
25	Arg		Ala	Asp	Tyr	Leu		Met	Leu	Leu	Phe			Ile	Cys	Ile	
		95					100					105	•				
														ATT			565
		lle	Thr	Gly	Leu			Asp	Met	Gln		Leu	Met	Ile	Pro		
20	110		=			115					120					125	
30														GAC			613
	TIE	Met	Ser	val		Tyr	Val	Trp	Ala		Leu	Asn	Arg	Asp		lle	
	O	mo 4	mmm	ma a	130	004		004		135			m 4 m	<b></b>	140	<b>700</b>	
														TTA			661
2 5	VAI	Ser	Pne	_	Phe	GLy	Thr	Arg		•	Ala	Cys	Tyr	Leu	Pro	Trp	
35	c m m	4 m O	0.00	145	mme		m 4 m		150			<b></b>		155	4.45	0.4.0	700
														ATC			709
	AST	TTE		GIY	rne	Asn	Tyr			Gly	Gly	ser		Ile	Asn	GIU	
			160					165					170				

	CTT ATT GGA AAT CTG GTT GGA CAT CTT	TAT TTT TTC CTA ATG TTC AGA 75	57
·	Leu Ile Gly Asn Leu Val Gly His Leu	Tyr Phe Phe Leu Met Phe Arg	
	175 180	185	
	TAC CCA ATG GAC TTG GGA GGA AGA AAT	TTT CTA TCC ACA CCT CAG TTT 80	)5
5	Tyr Pro Met Asp Leu Gly Gly Arg Asn	Phe Leu Ser Thr Pro Gln Phe	
	190 195	200 205	
•	TTG TAC CGC TGG CTG CCC AGT AGG AGA	GGA GGA GTA TCA GGA TTT GGT 85	53
	Leu Tyr Arg Trp Leu Pro Ser Arg Arg	Gly Gly Val Ser Gly Phe Gly	
	210	215 220	
10	GTG CCC CCT GCT AGC ATG AGG CGA GCT	GCT GAT CAG AAT GGC GGA GGC 90	1
	Val Pro Pro Ala Ser Met Arg Arg Ala	Ala Asp Gln Asn Gly Gly Gly	
	225 230	235	
	GGG AGA CAC AAC TGG GGC CAG GGC TTT	CGA CTT GGA GAC CAG TGAAGGG 95	0
	Gly Arg His Asn Trp Gly Gln Gly Phe	Arg Leu Gly Asp Gln	
15	240 245	. 250	
	GCGGCCTCGG GCAGCCGCTC CTCTCAAGCC ACA	TTTCCTC CCAGTGCTGG GTGCGCTTAA 101	0
	CAACTGCGTT CTGGCTAACA CTGTTGGACC TGAG	CCCACAC TGAATGTAGT CTTTCAGTAC 107	0
	GAGACAAAGT TTCTTAAATC CCGAAGAAAA ATA	FAAGTGT TCCACAAGTT TCACGATTCT 113	0
	CATTCAAGTC CTTACTGCTG TGAAGAACAA ATAC	CCAACTG TGCAAATTGC AAAACTGACT 119	0
20	ACATTTTTG GTGTCTTCTC TTCTCCCCTT TCCC	STCTGAA TAATGGGTTT TAGCGGGTCC 125	0
	TAGTCTGCTG GCATTGAGCT GGGGCTGGGT CACC	CAAACCC TTCCCAAAAG GACCCTTATC 131	0
	TCTTTCTTGC ACACATGCCT CTCTCCCACT TTTC	CCCAACC CCCACATTTG CAACTAGAAG 137	0
	AGGTTGCCCA TAAAATTGCT CTGCCCTTGA CAGG	GTTCTGT TATTTATTGA CTTTTGCCAA 143	0
	GGCTTGGTCA CAACAATCAT ATTCACGTAA TTTT	TCCCCCT TTGGTGGCAG AACTGTAGCA 149	0
25	ATAGGGGGAG AAGACAAGCA GCGGATGAAG CGTT	TTTCTCA GCTTTTGGAA TTGCTTCGAC 1550	0
	CTGACATCCG TTGTAACCGT TTGCCACTTC TTCA		0
	AGTCAGTGAG GGCCACAGAT TGGTATTAAT GAGA	,	0
	TTCCTGAGCT AAGTGATCAA GACTGTAGTG GAGT		0
	CCGTGGGGGA TGCAACCCCT TTGCGTTTCA TATG		0
30		•	0
	TCTTTGAGAG GCTCCTGGGC ATTGATTCCA TTTC		)
	GAGTAAAGGA GGAGAGACCC TCATACGCTA TTTA		)
	TTTTTTGGTC ATGTTTCAAT TAATTGTGAG GAAG		)
	ATTTTTAAA GCTAATGTAA GCACATCTAA GGGA		)
35			)
	AATCAGACCA GCTTAAATAC CCACACCTTT TTTT		)
	TTGGCTCATA ACCAAATAAA GTTTTTTGAA GGCC		•
	TTATGACGTT ATCTGAAAGC AGACTGTTAC CACC	ለርጥለጥጥ ርለርጥርርርጥርጥ ርልርልርጥጥቫርል ፡ 3331	١.

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•	ĢGCAACTAAA	AAGGCTTCAA	ACGTTTTGAT	CAGTTTCTTT	TCAGGAAACA	TTGTGCTCTA	2390
	ACAGTATGAC	TATTCTTTCC	CCCACTCTTA	AACAGTGTGA	TGTGTGTTAT	CCTAGGAAAT	2450
	GAGAGTTGGC	AAACAACTTC	TCATTTTGAA	TAGAGTTTGT	GTGTACCTCT	CCATATTTAA	2510
	TTTATATGAT	AAAATAGGTG	GGGAGAGTCT	GAACCTTAAC	TGTCATGTTT	TGTTGTTCAT	2570
5	CTGTGGCCAC	AATAAAGTTT	ACTTGTAAAA	TTTTAGAGGC	CATTACTCCA	ATTATGTTGC	2630
	ACGTACACTC	ATTGTACAGG	CGTGGAGACT	CATTGTATGT	ATAAGAATAT	TCTGACAGTG	2690
	AGTGACCCGG	AGTCTCTGGT	GTACCCTCTT	ACCAGTCAGC	TGCCTGCGAG	CAGTCATTTT	2750
	TTCCTAAAGG	TTTACAAGTA	TTTAGAACTC	TTCAGTTCAG	GGCAAAATGT	TCATGAAGTT	2810
	ATTCCTCTTA	AACATGGTTA	GGAAGCTGAT	GACGTTATTG	ATTTTGTCTG	GATTATGTTT	2870
10	CTGGAATAAT	TTTACCAAAA	CAAGCTATTT	GAGTTTTGAC	TTGACAAGGC	AAAACATGAC	2930
	AGTGGATTCT	CTTTACAAAT	TGAAAAAAA	AATCCTTATT	TTGTATAAAG	GACTTCCCTT	2990
:	TTTGTAAACT	AATCCTTTTT	ATTGGTAAAA	ATTGTAAATT	AAAATGTGCA	ACTTG	3045

- 15 (2) INFORMATION FOR SEQ ID NO: 43:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 653
    - (B) TYPE: Nucleic acid
    - (C) STRANDEDNESS: Double
- 20 (D) TOPOLOGY: Linear
  - (ii) SEQUENCE KIND: cDNA to mRNA
  - (vi) ORIGINAL SOURCE:
    - (A) ORGANISM: Homo sapiens
- 25 (B) CELL KIND: Epidermoid carcinoma
  - (C) CELL LINE: KB
  - (D) CLONE NAME: HP10389
  - (ix) SEQUENCE CHARACTERISTICS:
- 30 (A) CHARACTERIZATION CODE: CDS

35

- (B) EXISTENCE POSITION: 63.. 383
- (C) CHARACTERIZATION METHOD: E
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 43:

ATGACCTTCA CCGGGAGGCT GAGGTCGGAG TCCCGATTTT CTCCTGCTGC TGTGGCCCGG

AC ATG GCG ACT CCC GGC CCT GTG ATT CCG GAG GTC CCC TTT GAA CCA

Met Ala Thr Pro Gly Pro Val Ile Pro Glu Val Pro Phe Glu Pro

134

		1				5					10					15		
	TCG	AAG	CCT	CCA	GTC	ATT	GAG	GGG	CTG	AGC	CCC	ACT	GTT	TAC	AGG	AAT	3	155
	Ser	Lys	Pro	Pro	Val	Ile	Glu	Gly	Leu	Ser	Pro	Thr	Val	Tyr	Arg	Asn		
					20					25					30			
5	CCA	GAG	AGT	TTC	AAG	GAA	AAG	TTC	GTT	CGC	AAG	ACC	CGC	GAG	AAC	CCG	2	203
	Pro	Glu	Ser	Phe	Lys	G1u	Lys	Phe	Val	Arg	Lys	Thr	Arg	Glu	Asn	Pro		
				35					40					45				
	GTG	GTA	CCC	ATA	GGT	TGC	CTG	GCC	ACG	GCG	GCC	GCC	CTC	ACC	TAC	GGC	2	251
	Val	Val		Ile	Gly	Cys	Leu		Thr	Ala	Ala	Ala	Leu	Thr	Tyr	Gly		
10			50					55					60					
															ATG		2	99
	Leu		Ser	Pne	His	Arg		Asn	Ser	Gln	Arg		Gln	Leu	Met	Met		
	ccc	65	ccc	A TO CO	000	000	70	00m	mmo	400	omo.	75	000	4 M O	mmo	O TO C		, ,
15															TTG Leu		3	47
13	80	1111	VI R	116	VIG	85	GIII	GIY	rne	1111	90	VIR	WITE	116	Leu	95		
		CTG	GCT	GTC	АСТ		ATG	AAG	тст	CGA		ТААС	ecce/	, cc	STOTE	GCCTT	. 4	00
			Ala									******	,000			,00011	•	•
	,				100			-, -		105								
20	GAA	AGCTO	CCG C	CAGAA	ATGA	T TC	CAAA	ACC	AGG		AAC	CACT	rggco	CT A	ACCGI	GGGAC	4	60
	TTAC	CTCC	CTC C	CTCTC	CTTI	G AG	AGGC	CCAT	GTG	TCGC	TGG	GGAG	GAAG	TG A	ACCCI	TTGTG	5:	20
	TAAC	CTGTA	AAC C	GAAA	GTTI	T TI	CAAA	AATO	CTA	GATG	CTG	TTGT	TTGA	AT (	STTAC	ATACT	5	80
	TCTA	ATTTO	TG C	CACA	TCTC	c cc	TCCA	CTCC	CCI	GCTT	TAA'	AAAC	TCTA	LAA A	AATCC	ACTTG	6	40
	TAT	raat1	TTC A	AGT													6	53
25																		
	(2)		RMAI			•												
		t)	L) SE	•				ISTI	CS:									
		,			LENG													
30							clei											
							NESS			!								
		, ,					: Li											
		(1	i) S	EQUE	NCE	KIND	e cD	NA t	o mR	.NA							•	
35		( v	i) 0	PTCT	NAI	SUIDS	ርፑ.											
		( •	., o				: Ho	mo s	anie	ns								
										ance	r							
							_				_							

(D) CLONE NAME: HP10408

WO 98/55508

(ix) SEQUENCE CHARACTERISTICS:

(A) CHARACTERIZATION CODE: CDS

	(B)	EXISTENCE F	OSITION:	75 31	1		
	(C)	CHARACTERIZ	ATION ME	THOD: E			
5		1					
	(xi) SEQU	ENCE DESCRIP	TION: SE	Q ID NO:	44:		
	GTAGAAACAG GCCT	GTTAAG GAGAG	GCCAC CG	GGACTTCA	GTGTCTC	CTC CATC	CCAGGA 60
	GCGCAGTGGC CACT	ATG GGG TCT	GGG CTG	CCC CTT	GTC CTC	CTC TTG	ACC 110
10		Met Gly Ser	Gly Leu	Pro Leu	Val Leu	Leu Leu	Thr
		1	5			10	
	CTC CTT GGC AGC	TCA CAT GGA	ACA GGG	CCG GGT	ATG ACT	TTG CAA	CTG 158
	Leu Leu Gly Ser	Ser His Gly	Thr Gly	Pro Gly	Met Thr	Leu Gln	Leu
	15		20		25		
15	AAG CTG AAG GAG	TCT TTT CTG	ACA AAT	TCC TCC	TAT GAG	TCC AGC	TTC 206
	Lys Leu Lys Glu	Ser Phe Leu	Thr Asn	Ser Ser	Tyr Glu	Ser Ser	Phe
	30	35			40		•
	CTG GAA TTG CTT	GAA AAG CTC	TGC CTC	CTC CTC	CAT CTC	CCT TCA	GGG 254
	Leu Glu Leu Leu	Glu Lys Leu	Cys Leu	Leu Leu	His Leu	Pro Ser	Gly
20	45	50		55			60
	ACC AGC GTC ACC	CTC CAC CAT	GCA AGA	TCT CAA	CAC CAT	GTT GTC	TGC 302
	Thr Ser Val Thr	Leu His His	Ala Arg	Ser Gln	His His	Val Val	Cys
		65	•	70		75	
	AAC ACA TGACAGCO	CAT TGAAGCCT	GT GTCCT1	CTTG GCC	CCGGGCTT	TTGGGCCG	GG GA 360
25	Asn Thr						
	TGCAGGAGGC AGGCC	CCGAC CCTGT	CTTTC AGO	AGGCCCC	CACCCTCC	TG AGTGG	CAATA 420
	AATAAAATTC GGTAT	(GCTG					439
30							
	(2) INFORMATION	FOR SEQ ID 1	NO: 45:				
	(i) SEQUEN	NCE CHARACTE	RISTICS:				
	(A)	LENGTH: 1131	l				
	(B)	TYPE: Nuclei	ic acid				
35	(C)	STRANDEDNESS	S: Double				
	(D)	TOPOLOGY: Li	inear				
	(ii) SEQUE	ENCE KIND: cl	ONA to mR	NA	•		

		(,	V1)	ORIG	INAL	SOU	RCE:										
				(A)	ORG	ANIS	M: <i>H</i>	ото	sapi	ens							
				(B)	CEL	L KI	ND:	Stom	ach (	canc	er						
				(D)	CLO	NE N	AME:	HP1	0412								
5																	
		(:	ix)	SEQU	ENCE	CHAI	RACT	ERIS'	TICS	:					•		
				(A)	CHAI	RACT	ERIZ	ATIO	N CO	DE:	CDS						
				(B)	EXI	STEN	CE P	OSIT	ION:	56.	. 10	00	٠				
				(C)	CHA	RACT	ERIZ	ATIO	N ME	THOD	: E						
10																	
		(:	xi)	SEQU	ENCE	DES	CRIP'	TION	: SE	Q ID	NO:	45:					
	CTA:	rgag.	ATC (	CCGG	CCTC	AG G	GTGG	ACGC	A GT	GGTT	CTGC	ACT	GAGG	ccc ·	TCGT	C ATG	58
																Met	
15																1	
	GTG	GCG	CCT	GTG	TGG	TAC	TTG	GTA	GCG	GCG	GCT	CTG	CTA	GTC	GGC	TTT	106
	Val	Ala	Pro	Val	Trp	Tyr	Leu	Val	Ala	Ala	Ala	Leu	Leu	Val	Gly	Phe	
				5					10					15			
	ATC	CTC	TTC	CTG	ACT	CGC	AGC	CGG	GGC	CGG	GCG	GCA	TCA	GCC	GGC	CAA	154
20	Ile	Leu	Phe	Leu	Thr	Arg	Ser	Arg	Gly	Arg	Ala	Ala	Ser	Ala	Gly	Gln	
			20					25					30				
	GAG	CCA	CTG	CAC	AAT	GAG	GAG	CTG	GCA	GGA	GCA	GGC	CGG	GTG	GCC	CAG	202
	Glu	Pro	Leu	His	Asn	Glu	Glu	Leu	Ala	Gly	Ala	Gly	Arg	Val	Ala	Gln	
		35					40					45					
25	CCT	GGG	CCC	CTG	GAG	CCT	GAG	GAG	CCG	AGA	GCT	GGA	GGC	AGG	CCT	CGG	250
	Pro	Gly	Pro	Leu	Glu	Pro	Glu	Glu	Pro	Arg	Ala	Gly	Gly	Arg	Pro	Arg	
	50					55					60					65	
				GAC													298
	Arg	Arg	Arg	Asp	Leu	Gly	Ser	Arg	Leu	Gln	Ala	Gln	Arg	Arg		Gln	
30					70					75					80		
				TGG													346
	Arg	Val	Ala	Trp	Ala	Glu	Ala	Asp	Glu	Asn	Glu	Glu	Glu		Val	Ile	
				85					90					95			
				GAG													394
35	Leu	Ala		Glu	Glu	Glu	G1y		Glu	Lys	Pro	Ala		Thr	His	Leu	
			100					105					110			0//	
				ATT													442
	Ser	Gly	Lys	Ile	Gly	Ala	Lys	Lys	Leu	Arg	Lys	Leu	Glu	Glu	Lys	Gin	

		115				_	120					125	1				
	GCG	CGA	AAG	GCC	CAG	CGT	GAG	GCA	GAG	GAG	GCT	GAA	CGT	GAG	GAG	CGG	490
	Ala	Arg	Lys	Ala	G1n	Arg	Glu	Ala	Glu	Glu	Ala	Glu	Arg	Glu	Glu	Arg	•
	130					135					140					145	
5	AAA	CGA	CTC	GAG	TCC	CAG	CGC	GAA	GCT	GAG	TGG	AAG	AAG	GAG	GAG	GAG	538
	Lys	Arg	Leu	Glu	Ser	Gln	Arg	Glu	Ala	Glu	Trp	Lys	Lys	Glu	Glu	Glu	
					150					155					160	)	
	CGG	CTT	CGC	CTG	GAG	GAG	GAG	CAG	AAG	GAG	GAG	GAG	GAG	AGG	AAG	GCC	586
	Arg	Leu	Arg	Leu	Glu	Glu	Glu	Gln	Lys	Glu	Glu	Glu	Glu	Arg	Lys	Ala	
10				165					170					175			
	CGC	GAG	GAG	CAG	GCC	CAG	CGG	GAG	CAT	GAG	GAG	TAC	CTG	AAA	CTG	AAG	634
	Arg	Glu	Glu	Gln	Ala	Gln	Arg	Glu	His	Glu	Glu	Tyr	Leu	Lys	Leu	Lys	
			180					185					190				
	GAG	GCC	TTT	GTG	GTG	GAG	GAG	GAA	GGC	GTA	GGA	GAG	ACC	ATG	ACT	GAG	682
15	Glu	Ala	Phe	Val	Val	Glu	Glu	Glu	Gly	Val	Gly	Glu	Thr	Met	Thr	Glu	
		195					200					205					•
	GAA	CAG	TCC	CAG	AGC	TTC	CTG	ACA	GAG	TTC	ATC	AAC	TAC	ATC	AAG	CAG	730
	Glu	Gln	Ser	G1n	Ser	Phe	Leu	Thr	Glu	Phe	Ile	Asn	Tyr	Ile	Lys	Gln	
	210					215					220					225	
20	TCC	AAG	GTT	GTG	CTC	TTG	GAA	GAC	CTG	GCT	TCC	CAG	GTG	GGC	CTA	CGC	778
	Ser	Lys	Val	Val	Leu	Leu	Glu	Asp	Leu	Ala	Ser	Gln	Val	Gly	Leu	Arg	
					230					235					240		
	ACT	CAG	GAC	ACC	ATA	AAT	CGC	ATC	CAG	GAC	CTG	CTG	GCT	GAG	GGG	ACT	826
	Thr	Gln	Asp	Thr	Ile	Asn	Arg	Ile	Gln	Asp	Leu	Leu	Ala	Glu	Gly	Thr	
25				245					250					255			
	ATA	ACA	GGT	GTG	TTA	GAC	GAC	CGG	GGC	AAG	TTC	ATC	TAC	ATA	ACC	CCA	874
	Ile	Thr	Gly	Val	Ile	Asp	Asp	Arg	Gly	Lys	Phe	Ile	Tyr	Ile	Thr	Pro	
			260					265					270				
							GCC .										922
30	G1u		Leu	Ala	Ala	Val	Ala	Asn	Phe	Ile	Arg	Gln	Arg	Gly	Arg	Val	
		275					280					285					
							CAA										970
		Ile	Ala	Glu	Leu	Ala	Gln	Ala	Ser	Asn	Ser	Leu	Ile	Ala	Trp	Gly	
	290					295					300				,	305	
35							GCC			TGAC	CCCA	GT C	CTTC	CCTC	T TO	G.	1020
	Arg	Glu	Ser	Pro		G1n	Ala	Pro	Ala								
			_		310												
	ACTO	AGAG	TT G	GTGI	GGCC	T AC	CTGG	CTAT	ACA	TCTT	CAT	CCCT	cccc	AC C	ATCC	TGGGG	1080

138

1131

AAGTGATGGT GTGGCCAGGC AGTTATAGAT TAAAGGCCTG TGAGTACTGC T

	(2) INFORMATI	ON FOR SEQ II	NO: 46:												
5	(i) SEQ	UENCE CHARACT	ERISTICS:												
	(.	A) LENGTH: 18	375												
	(	B) TYPE: Nucl	eic acid												
	(	C) STRANDEDNE	SS: Doubl	е											
	(	D) TOPOLOGY:	Linear												
10	(ii) SE	QUENCE KIND:	cDNA to m	RNA											
						•									
	(vi) OR	IGINAL SOURCE	:												
	(,	A) ORGANISM:	Homo sapi	ens											
	. (3	B) CELL KIND:	Stomach	cancer											
15	(1	D) CLONE NAME	: HP10413												
	(iv) SE	TIPNOS CHADAC	TED T CTTOC												
	(ix) SEQUENCE CHARACTERISTICS:  (A) CHARACTERIZATION CODE: CDS														
	(B) EXISTENCE POSITION: 79 666														
20	•	C) CHARACTERI			ь										
	(,	) ommercible	EXITON THE	Inob. E											
	(xi) SEC	QUENCE DESCRI	PTION: SE	Q ID NO:	46:										
	·														
	CTCGCTCGCT CAC	AGGGAGG AGAA	AGTGGC GA	GTTCCGGA	TCCCTGC	CTA GCGC	GCCCA	60							
25	ACCTTTACTC CAC	AGATC ATG GC	T GCC GAG	GAT GTG	GTG GCG	ACT GGC	GCC	111							
		Met Al	a Ala Glu	Asp Val	Val Ala	Thr Gly	Ala								
		1		5		10									
	GAC CCA AGC GA	AT CTG GAG AG	c GGC GGG	CTG CTG	CAT GAG	ATT TTC	ACG	159							
	Asp Pro Ser As	p Leu Glu Se	r Gly Gly	Leu Leu	His Glu	Ile Phe	Thr								
30	· 1	L <b>5</b>	20			25									
	TCG CCG CTC AA	C CTG CTG CT	G CTT GGC	CTC TGC	ATC TTC	CTG CTC	TAC	207							
	Ser Pro Leu As	n Leu Leu Le	u Leu Gly	Leu Cys	Ile Phe	Leu Leu	Tyr								
	30		35		40										
	AAG ATC GTG CG	C GGG GAC CA	G CCG GCG	GCC AGC	GGC GAC	AGC GAC	GAC	255							
35	Lys Ile Val Ar	g Gly Asp Gl	n Pro Ala	Ala Ser	Gly Asp	Ser Asp	Asp								
	45	50			55										
	GAC GAG CCG CC	C CCT CTG CC	C CGC CTC	AAG CGG	CGC GAC	TTC ACC	CCC	303							

Asp Glu Pro Pro Pro Leu Pro Arg Leu Lys Arg Arg Asp Phe Thr Pro

	· 60					65					70					75	
	GCC	GAG	CTG	CGG	CGC	TTC	GAC	GGC	GTC	CAG	GAC	CCG	CGC	ATA	CTC	ATG	351
	Ala	G1u	Leu	Arg	Arg	Phe	Asp	Gly	Val	Gln	Asp	Pro	Arg	Ile	Leu	Met	
					80					85					90		
5	GCC	ATC	AAC	GGC	AAG	GTG	TTC	GAT	GTG	ACC	AAA	GGC	CGC	AAA	TTC	TAC	399
	Ala	Ile	Asn	Gly	Lys	Val	Phe	Asp	Val	Thr	Lys	Gly	Arg	Lys	Phe	Tyr	
				95					100					105			
	GGG	ССС	GAG	GGG	CCG	TAT	GGG	GTC	TTT	GCT	GGA	AGA	GAT	GCA	TCC	AGG	447
	Gly	Pro	Glu	Gly	Pro	Tyr	Gly	Val	Phe	Ala	Gly	Arg	Asp	Ala	Ser	Arg	
10			110					115					120				
	GGC	CTT	GCC	ACA	TTT	TGC	CTG	GAT	AAG	GAA	GCA	CTG	AAG	GAT	GAG	TAC	495
	Gly	Leu	Ala	Thr	Phe	Cys	Leu	Asp	Lys	Glu	Ala	Leu	Lys	Asp	Glu	Tyr	
		125					130					135					
	GAT	GAC	CTT	TCT	GAC	CTC	ACT	GCT	GCC	CAG	CAG	GAG	ACT	CTG	AGT	GAC	543
15	Asp	Asp	Leu	Ser	Asp	Leu	Thr	Ala	Ala	Gln	Gln	Glu	Thr	Leu	Ser	Asp	
	140					145					150					155	
	TGG	GAG	TCT	CAG	TTC	ACT	TTC	AAG	TAT	CAT	CAC	GTG	GGC	AAA	CTG	CTG	591
	Trp	Glu	Ser	Gln	Phe	Thr	Phe	Lys	Tyr	His	His	Val	Gly	Lys	Leu	Leu	
					160					165					170		
20	AAG	GAG	GGG	GAG	GAG	ccc	ACT	GTG	TAC	TCA	GAT	GAG	GAA	GAA	CCA	AAA	639
	Lys	Glu	Gly	Glu	Glu	Pro	Thr	Val	Tyr	Ser	Asp	Glu	Glu	Glu	Pro	Lys	
				175					180					185			
	GAT	GAG	AGT	GCC	CGG	AAA	TAA	GAT	TAAA	GCAI	TC A	GTGG	AAGI	A TA	ATCTA	T	690
	Asp	Glu	Ser	Ala	Arg	Lys	Asn	Asp									
25			190					195									
	TTTT	GTA1	rtt 1	TGCAA	AATC	A TI	TGTA	ACAG	TCC	ACTO	TGT	CTTI	'AAAA	CA I	CAGTG	ATTAC	750
	AATA	TTTA	AGA A	AAGTI	TTGA	G CA	CTTG	CTAT	AAG	TTTI	TTA	TAAC	ATCA	CT A	AGTGA	CACTA	810
	ATAA	AATI	C AA1	CTTCI	TAGA	A TG	CATG	ATGT	GTI	TGTG	TGT	CACA	AATO	CA	AAAG	TGAAC	870
	TGCA	GTGC	CTG 1	ATAAT	CACA	T GI	'TAAT	ACTG	TTT	TTCI	TCT	ATCI	GTAG	TT A	AGTAC	AGGAT	930
30	GAAT	'ATTA	TAA	STGTT	TTTC	C TG	AGAG	ACAA	GGA	AGAC	TTG	GGTA	TTTC	CC A	LAAAC	AGGTA	990
	AAAA	TCTI	AA7	ATGTG	CACC	A AG	AGCA	AAGG	ATC	AACT	TTT	AGTO	ATGA	TG I	TCTG	TAAAG	1050
	ACAA	CAAA	ATC (	CTTI	TTTT	T TC	TCAA	TTGA	CTT	AACT	GCA	TGAT	TTCI	GT I	TATT	CTACC	1110
	TCTA	AAGC	CAA A	ATCTG	CAGT	G TI	'CCAA	AGAC	TTT	GGTA	TGG	ATTA	AGCG	CT G	TCCA	GTAAC	1170
	AAAA	TGAA	AAT (	CTCAA	AACA	G AG	CTCA	GCTG	CAA	AAAA	GCA	TATI	TTCI	GT G	TTTC	TGGAC	1230
35	TGCA	CTGI	TG 1	CCTI	'GCCC	T CA	CATA	GACA	CTC	AGAC	ACC	CTCA	.CAAA	CA C	CAGTA	GTCTA	1290
	TAGT	TAGO	AT 1	AAAA7	TAGG	A TO	TGAA	CATT	CAA	AAGA	AAG	CTTT	GGAA	AA A	AAGA	GCTGG	1350
	CTGG	CCTA	AAA A	ACCI	TAAA'	A TA	TGAI	'GAAG	ATT	GTAG	GAC	TGTC	TTCC	CA A	GCCC	CATGT	1410
	TCAT	GGTC	GG G	CAAT	GGTT	A TT	TGGT	TATT	TTA	CTCA	ATT	GGTT	ACTO	TC A	TTTG	AAATG	1470

WO 98/55508

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AGGGAGGGAC ATACAGAATA GGAACAGGTG TTTGCTCTCC TAAGAGCCTT CATGCACACC

PCT/JP98/02445

	CCTGAACCAC GAGGAAACAG TACAGTCGCT AGTCAAGTGG TTTTTAAAGT AAAGTATATT	1590
	CATAAGGTAA CAGTTATTCT GTTGTTATAA AACTATACCC ACTGCAAAAG TAGTAGTCAA	1650
	GTGTCTAGGT CTTTGATATT GCTCTTTTGG TTAACACTAA GCTTAAGTAG ACTATACAGT	1710
5	TGTATGAATT TGTAAAAGTA TATGAACACC TAGTGAGATT TCAAACTTGT AATTGTGGTT	1770
	AAATAGTCAT TGTATTTTCT TGTGAACTGT GTTTTATGAT TTTACCTCAA ATCAGAAAAC	1830
	AAAATGATGT GCTTTGGTCA GTTAATAAAA ATGGTTTTAC CCACT	1875
,		
. 10	(2) INFORMATION FOR SEQ ID NO: 47:	
	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 1563	
	(B) TYPE: Nucleic acid	
	(C) STRANDEDNESS: Double	
15	(D) TOPOLOGY: Linear	
	(ii) SEQUENCE KIND: cDNA to mRNA	
	(vi) ORIGINAL SOURCE:	
	(A) ORGANISM: Homo sapiens	
20	(B) CELL KIND: Stomach cancer	
•	(D) CLONE NAME: HP10415	
	(ix) SEQUENCE CHARACTERISTICS:	
	(A) CHARACTERIZATION CODE: CDS	
25	(B) EXISTENCE POSITION: 72 1460	
	(C) CHARACTERIZATION METHOD: E	
	<b>(3, 3, 3, 3, 3, 3, 3, 3, 3, 3, 3, 3, 3, 3</b>	
•	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 47:	
30	AAATTGGGCC AGGCTGAGGC GCTGCTGCTG GAGCGGCCGA TCCGAGACGT GGCTCCCTGG	60
	GCGGCAGAAC C ATG TTG GAC TTC GCG ATC TTC GCC GTT ACC TTC TTG CTG	110
	Met Leu Asp Phe Ala Ile Phe Ala Val Thr Phe Leu Leu	
	1 5 10	
	GCG TTG GTG GGA GCC GTG CTC TAC CTC TAT CCG GCT TCC AGA CAA GCT	158
35	Ala Leu Val Gly Ala Val Leu Tyr Leu Tyr Pro Ala Ser Arg Gln Ala	
	15 20 25	
	GCA GGA ATT CCA GGG ATT ACT CCA ACT GAA GAA AAA GAT GGT AAT CTT	206
	Ala Gly Ile Pro Gly Ile Thr Pro Thr Glu Glu Lys Asp Gly Asn Leu	

	30					35					40					45	
	CCA	GAT	ATT	GTG	AAT	AGT	GGA	AGT	TTG	CAT	GAG	TTC	CTG	GTT	AAT	TTG	254
	Pro	Asp	Ile	Val	Asn	Ser	Gly	Ser	Leu	His	Glu	Phe	Leu	Val	Asn	Leu	
					50					55					60		
5	CAT	GAG	AGA	TAT	GGG	CCT	GTG	GTC	TCC	TTC	TGG	TTT	GGC	AGG	CGC	CTC	302
	His	Glu	Arg	Tyr	Gly	Pro	Val	Val	Ser	Phe	Trp	Phe	Gly	Arg	Arg	Leu	
				65					70					75			
	GTG	GTT	AGT	TTG	GGC	ACT	GTT	GAT	GTA	CTG	AAG	CAG	CAT	ATC	AAT	CCC	350
	Val	Val	Ser	Leu	Gly	Thr	Val	Asp	Val	Leu	Lys	Gln	His	Ile	Asn	Pro	
10			80					85					90				
	AAT	AAG	ACA	TTG	GAC	CCT	TTT	GAA	ACC	ATG	CTG	AAG	TCA	TTA	TTA	AGG	398
	Asn	Lys	Thr	Leu	Asp	Pro	Phe	Glu	Thr	Met	Leu	Lys	Ser	Leu	Leu	Arg	
		95					100					105					
	TAT	CAA	TCT	GGT	GGT	GGC	AGT	GTG	AGT	GAA	AAC	CAC	ATG	AGG	AAA	AAA	446
15	Tyr	Gln	Ser	Gly	Gly	Gly	Ser	Val	Ser	Glu	Asn	His	Met	Arg	Lys	Lys	
	110					115					120					125	
	TTG	TAT	GAA	AAT	GGT	GTG	ACT	GAT	TCT	CTG	AAG	AGT	AAC	TTT	GCC	CTC	494
	Leu	Tyr	Glu	Asn	Gly	Val	Thr	Asp	Ser	Leu	Lys	Ser	Asn	Phe	Ala	Leu	
					130					135					140		
20	CTC	CTA	AAG	CTT	TCA	GAA	GAA	TTA	TTA	GAT	AAA	TGG	CTC	TCC	TAC	CCA	542
	Leu	Leu	Lys	Leu	Ser	Glu	Glu	Leu	Leu	Asp	Lys	Trp	Leu	Ser	Tyr	Pro	
				145					150					155			
	GAG	ACC	CAG	CAC	GTG	CCC	CTC	AGC	CAG	CAT	ATG	CTT	GGT	TTT	GCT	ATG	590
	Glu	Thr	Gln	His	Val	Pro	Leu	Ser	Gln	His	Met	Leu	Gly	Phe	Ala	Met	
25			160					165					170				
	AAG	TCT	GTT	ACA	CAG	ATG	GTA	ATG	GGT	AGT	ACA	TTT	GAA	GAT	GAT	CAG	638
	Lys	Ser	Val	Thr	Gln	Met	Val	Met	Gly	Ser	Thr	Phe	Glu	Asp	Asp	Gln	
		175					180					185					
													TGG				686
30	Glu	Val	Ile	Arg	Phe	Gln	Lys	Asn	His	Gly	Thr	Val	Trp	Ser	Glu		
	190					195					200					205	
													ATG				734
	Gly	Lys	Gly	Phe	Leu	Asp	Gly	Ser	Leu	Asp	Lys	Asn	Met	Thr	Arg	Lys	
					210					215					220		
35													GTT				782
	Lys	Gln	Tyr		Asp	Ala	Leu	Met		Leu	Glu	Ser	Val		Arg	Asn	
				225					230					235			
	ል ጥር	ΔΤΔ	A A A	CAA	CCA	A A A	CCA	ACC	AAC	ጥጥር	ACT	CAA	CAT	ΑTΤ	TTC	ATT	830

	Ile	Ile		Glu	Arg	Lys	Gly	_	Asn	Phe	Ser	Gln		Ile	Phe	Ile	,
			240					245					250				
																GAC	878
	Asp	Ser	Leu	Val	Gln	Gly	Asn	Leu	Asn	Asp	Gln	Gln	Ile	Leu	Glu	Asp	
5		255					260					265					
	AGT	ATG	ATA	TTT	TCT	CTG	GCC	AGT	TGC	ATA	ATA	ACT	GCA	AAA	TTG	TGT	926
	Ser	Met	Ile	Phe	Ser	Leu	Ala	Ser	Cys	Ile	Ile	Thr	Ala	Lys	Leu	Cys	
	270					275					280					285	
	ACC	TGG	GCA	ATC	TGT	TTT	TTA	ACC	ACC	TCT	GAA	GAA	GTT	CAA	AAA	AAA	974
10	Thr	Trp	Ala	Ile	Cys	Phe	Leu	Thr	Thr	Ser	Glu	Glu	Val	Gln	Lys	Lys	
			•		290					295					300		
•	TTA	TAT	GAA	GAG	ATA	AAC	CAA	GTT	TTT	GGA	AAT	GGT	CCT	GTT	ACT	CCA	1022
	Leu	Tyr	Glu	Glu	Ile	Asn	Gln	Val	Phe	Gly	Asn	Gly	Pro	Val	Thr	Pro	
				305					310					315			
15															GAA		1070
	Glu	Lys	Ile	Glu	Gln	Leu	Arg	Tyr	Cys	Gln	His	Val	Leu	Cys	Glu	Thr	
			320					325					330				
	GTT	CGA	ACT	GCC	AAA	CTG	ACT	CCA	GTT	TCT	GCC	CAG	CTT	CAA	GAT	ATT	1118
	Val	Arg	Thr	Ala	Lys	Leu	Thr	Pro	Val	Ser	Ala	G1n	Leu	Gln	Asp	Ile	
20		335					340					345					
	GAA	GGA	AAA	ATT	GAC	CGA	TTT	ATT	ATT	CCT	AGA	GAG	ACC	CTC	GTC	CTT	1166
		Gly	Lys	Ile	Asp	Arg	Phe	Ile	Ile	Pro	Arg	Glu	Thr	Leu	Val		
	350					355					360					365	
															TCT		1214
25	Tyr	Ala	Leu	G1y	Val	Val	Leu	Gln	Asp	Pro	Asn	Thr	Trp	Pro	Ser	Pro	
					370					375					380		
															AAA		1262
	His	Lys	Phe	-	Pro	Asp	Arg	Phe	-	Asp	Glu	Leu	Val		Lys	Thr	
20				385					390					395			
30															TTG		1310
	rne	ser		Leu	GIÀ	Pne	ser	-	Thr	GIn	GIU	Cys		GIU	Leu	Arg	
			400	4 ma		4.00		405					410	a.m.a			
															AAG		1358
2 =	rne		ıyr	met	Val	rnr		VAI	Leu	Leu			ren	val	Lys	Arg	
35	CTC.	415	Cm 4	O m m	mo=	C MC	420	004	0.0	0 m.c		425	۸۵.	440	m v w	C A A	1400
															TAT		1406
		urs	ren	ren	ser		GIU	GIÀ	GIN	val		GIU	ınr	гàг	Tyr		
	430					435					440		•			445	

	CTG GTA ACA TCA TCA AGG GAA GAA GCT TGG ATC ACT GTC TCA AAG AGA	145
	Leu Val Thr Ser Ser Arg Glu Glu Ala Trp Ile Thr Val Ser Lys Arg	
	450 455 460	
	TAT TAAAATTTTA TACATTTAAA ATCATTGTTA AATTGATTGA GGAAAACAAC CAT	151
5	Tyr	
	TTAAAAAAA TCTATGTTGA ATCCTTTTAT AAACCAGTAT CACTTTGTAA TAT	156
1 0	(a) Typopy( many non-one to ye	
10	(2) INFORMATION FOR SEQ ID NO: 48:	
	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 2030	
	(B) TYPE: Nucleic acid (C) STRANDEDNESS: Double	
15	(D) TOPOLOGY: Linear	
	(ii) SEQUENCE KIND: cDNA to mRNA	
	(vi) ORIGINAL SOURCE:	
	(A) ORGANISM: Homo sapiens	
20	(B) CELL KIND: Stomach cancer	
	(D) CLONE NAME: HP10419	
	(ix) SEQUENCE CHARACTERISTICS:	
	(A) CHARACTERIZATION CODE: CDS	
25	(B) EXISTENCE POSITION: 171 914	
	(C) CHARACTERIZATION METHOD: E	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 48:	
30	CATTIGGGGT TICGGTTCCC CCCCTTCCCC TICCCCGGGG TCTGGGGGTG ACATTGCACC	60
-	GCGCCCTCG TGGGGTCGCG TTGCCACCC ACGCGGACTC CCCAGCTGGC GCGCCCCTCC	120
	CATTTGCCTG TCCTGGTCAG GCCCCCACCC CCCTTCCCAC CTGACCAGCC ATG GGG	176
	Met Gly	
	1	
35	GCT GCG GTG TTT TTC GGC TGC ACT TTC GTC GCG TTC GGC CCG GCC TTC	224
	Ala Ala Val Phe Phe Gly Cys Thr Phe Val Ala Phe Gly Pro Ala Phe	
	5 10 15	
	GCG CTT TTC TTG ATC ACT GTG GCT GGG GAC CCG CTT CGC GTT ATC ATC	272

	Ala	Leu	Phe	Leu	Ile	Thr	Val	Ala	Gly	Asp	Pro	Leu	Arg	Val	Ile	Ile	
		20					25					30					
	CTG	GTC	GCA	GGG	GCA	TTT	TTC	TGG	CTG	GTC	TCC	CTG	CTC	CTG	GCC	TCT	320
	Leu	Val	Ala	Gly	Ala	Phe	Phe	Trp	Leu	Val	Ser	Leu	Leu	Leu	Ala	Ser	
5	35					40					45					50	
	GTG	GTC	TGG	TTC	ATC	TTG	GTC	CAT	GTG	ACC	GAC	CGG	TCA	GAT	GCC	CGG	368
	Val	Val	Trp	Phe	Ile	Leu	Val	His	Va1	Thr	Asp	Arg	Ser	Asp	Ala	Arg	
					55					60					65		
	CTC	CAG	TAC	GGC	CTC	CTG	ATT	TTT	GGT	GCT	GCT	GTC	TCT	GTC	CTT	CTA	416
10	Leu	Gln	Tyr	G1y	Leu	Leu	Ile	Phe	Gly	Ala	Ala	Val	Ser	Val	Leu	Leu	
				70					75					80			
	CAG	GAG	GTG	TTC	CGC	TTT	GCC	TAC	TAC	AAG	CTG	CTT	AAG	AAG	GCA	GAT	464
	Gln	Glu	Val	Phe	Arg	Phe	Ala	Tyr	Tyr	Lys	Leu	Leu	Lys	Lys	Ala	Asp	
			85					90					95				
15	GAG	GGG	TTA	GCA	TCG	CTG	AGT	GAG	GAC	GGA	AGA	TCA	ccc	ATC	TCC	ATC	512
	Glu	Gly	Leu	Ala	Ser	Leu	Ser	Glu	Asp	Gly	Arg	Ser	Pro	Ile	Ser	Ile	
		100					105					110			•		
	CGC	CAG	ATG	GCC	TAT	GTT	TCT	GGT	CTC	TCC	TTC	GGT	ATC	ATC	AGT	GGT	560
	Arg	Gln	Met	Ala	Tyr	Va1	Ser	Gly	Leu	Ser	Phe	Gly	Ile	Ile	Ser	Gly	
20	115					120					125					130	
	GTC	TTC	TCT	GTT	ATC	TAA	ATT	TTG	GCT	GAT	GCA	CTT	GGG	CCA	GGT	GTG	608
,	Val	Phe	Ser	Val	Ile	Asn	Ile	Leu	Ala	Asp	Ala	Leu	Gly	Pro	Gly	Val	
					135					140					145		
	GTT	GGG	ATC	CAT	GGA	GAC	TCA	CCC	TAT	TAC	TTC	CTG	ACT	TCA	GCC	TTT	656
25	Val	Gly	Ile	His	Gly	Asp	Ser	Pro	Tyr	Tyr	Phe	Leu	Thr	Ser	Ala	Phe	
				150					155					160			
	CTG	ACA	GCA	GCC	ATT	ATC	CTG	CTC	CAT	ACC	TTT	TGG	GGA	GTT	GTG	TTC	704
	Leu	Thr	Ala	Ala	Ile	Ile	Leu	Leu	His	Thr	Phe	Trp	Gly	Val	Val	Phe	
			165					170					175				
30	TTT	GAT	GCC	TGT	GAG	AGG	AGA	CGG	TAC	TGG	GCT	TTG	GGC	CTG	GTG	GTT	752
	Phe	Asp	Ala	Cys	Glu	Arg	Arg	Arg	Tyr	Trp	Ala	Leu	Gly	Leu	Val	Val	
		180					185					190					
	GGG	AGT	CAC	CTA	CTG	ACA	TCG	GGA	CTG	ACA	TTC	CTG	AAC	CCC	TGG	TAT	800
	Gly	Ser	His	Leu	Leu	Thr	Ser	Gly	Leu	Thr	Phe	Leu	Asn	Pro	Trp	Tyr	
35	195					200					205					210	
	GAG	GCC	AGC	CTG	CTG	CCC	ATC	TAT	GCA	GTC	ACT	GTT	TCC	ATG	GGG	CTC	848
	Glu	Ala	Ser	Leu	Leu	Pro	Ile	Tyr	Ala	Val	Thr	Val	Ser	Met	Gly	Leu	
					215					220					225		

## 145

	TGG GCC TTC ATC ACA G	CT GGA GGG	TCC CTC CGA	AGT ATT CAG CGC AGC	896
	Trp Ala Phe Ile Thr A	la Gly Gly	Ser Leu Arg	Ser Ile Gln Arg Ser	
	230		235	240	
	CTC TTG TGT AAG GAC T	GACTACCTG G	ACTGATCGC C	TGACAGATC CCACCTGCC	950
5	Leu Leu Cys Lys Asp				
	245				
	TGTCCACTGC CCATGACTGA	GCCCAGCCCC	AGCCCGGGTC	CATTGCCCAC ATTCTCTGTC	1010
	TCCTTCTCGT CGGTCTACCC	CACTACCTCC	AGGGTTTTGC	TTTGTCCTTT TGTGACCGTT	1070
	AGTCTCTAAG CTTTACCAGG	AGCAGCCTGG	GTTCAGCCAG	TCAGTGACTG GTGGGTTTGA	1130
10	ATCTGCACTT ATCCCCACCA	CCTGGGGACC	CCCTTGTTGT	GTCCAGGACT CCCCCTGTGT	1190
	CAGTGCTCTG CTCTCACCCT	GCCCAAGACT	CACCTCCCTT	CCCCTCTGCA GGCCGACGGC	1250
	AGGAGGACAG TCGGGTGATG	GTGTATTCTG	CCCTGCGCAT	CCCACCCGAG GACTGAGGGA	1310
	ACCTAGGGG GACCCCTGGG	CCTGGGGTGC	CCTCCTGATG	TCCTCGCCCT GTATTTCTCC	1370
	ATCTCCAGTT CTGGACAGTG	CAGGTTGCCA	AGAAAAGGGA	CCTAGTTTAG CCATTGCCCT	1430
<b>L</b> 5	GGAGATGAAA TTAATGGAGG	CTCAAGGATA	GATGAGCTCT	GAGTTTCTCA GTACTCCCTC	1490
	AAGACTGGAC ATCTTGGTCT	TTTTCTCAGG	CCTGAGGGGG	AACCATTTTT GGTGTGATAA	1550
	ATACCCTAAA CTGCCTTTTT	TTCTTTTTTG	AGGTGGGGG	AGGGAGGAGG TATATTGGAA	1610
	CTCTTCTAAC CTCCTTGGGC	TATATTTCT	CTCCTCGAGT	TGCTCCTCAT GGCTGGGCTC	1670
	ATTTCGGTCC CTTTCTCCTT	GGTCCCAGAC	CTTGGGGGAA	AGGAAGGAAG TGCATGTTTG	1730
20	GGAACTGGCA TTACTGGAAC	TAATGGTTTT	AACCTCCTTA	ACCACCAGCA TCCCTCCTCT	1790
	CCCCAAGGTG AAGTGGAGGG	TGCTGTGGTG	AGCTGGCCAC	TCCAGAGCTG CAGTGCCACT	1850
	GGAGGAGTCA GACTACCATG	ACATCGTAGG	GAAGGAGGG	AGATTTTTTT GTAGTTTTTA	1910
•	ATTGGGGTGT GGGAGGGGCG	GGGAGGTTTT	CTATAAACTG	TATCATTTTC TGCTGAGGGT	1970
	GGAGTGTCCC ATCCTTTTAA	TCAAGGTGAT	TGTGATTTTG	ACTAATAAAA AAGAATTTGT	2030
25	·				

(2) INFORMATION FOR SEQ ID NO: 49:

- (1) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 493
  - (B) TYPE: Nucleic acid
    - (C) STRANDEDNESS: Double
    - (D) TOPOLOGY: Linear
- (ii) SEQUENCE KIND: cDNA to mRNA
- 35 (vi) ORIGINAL SOURCE:

- (A) ORGANISM: Homo sapiens
- (B) CELL KIND: Stomach cancer
- (D) CLONE NAME: HP10424

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(ix) SEQUENCE CHARACTERISTICS:

				(A)	CHA	RACT	ERIZ	ATIO	N CO	DE:	CDS						
				(B)	EXI	STEN	CE P	OSIT	ION:	98.	. 43	9		:			
				(C)	CHA	RACT	ERIZ	ATIO	N ME	THOD	: E						
5																	
		(	xi)	SEQU	ENCE	DES	CRIP	TION	: SE	Q ID	NO:	49:					
										•							
	AAA	GTTT	ccc .	AAAT	CCAG	GC G	GCTA	GAGG	C CC	ACTG	CTTC	CCA	ACTA	CCA	GCTG	AGGGGG	60
	TCC	GTCC	CGA (	GAAG	GGAG.	AA G	AGGC	CGAA	G AĞ	GAAA	C AT	G AA	C TT	C TA	T TT.	A CTC	115
10																u Leu	
												1				5	
	CTA	GCG	AGC	AGC	ATT	CTG	TGT	GCC	TTG	ATT	GTC	TTC	TGG	AAA	TAT	CGC	163
															Tyr		
				10			-,-		15					20	- ) -		
15	CGC	ттт	CAG		AAC	ACT	GGC	GAA		TCA	TCA	ΔΑΤ	тса		GCT	СТТ	211
															Ala		~
	8		25	6			01)	30	1100	Jer	561	non	35	****	2114	Dea	
	GCA	СТА		AGA	ccc	ጥርጥ	ጥረጥ		ccc	m m A	ል ጥጥ	A A C		ል ል ጥ	ACA	CAC	259
	•														Thr		239
20	N10	40	Va1	nr g	110	261		Ser	Gry	Leu	116		Ser	VPII	1111	veh	
20	440		C TO TO	CC 4	C TIC	m A C	45	0.00	mom.		0.48	50		4 A M	4 A M	mmo	207
															AAT		307
		ASI	ren	AIA	VAI		Asp	Leu	Ser	Arg	_	11e	Leu	Asn	Asn.		
	55					60					65					70	
^-															AGT		355
25	Pro	His	Ser	Ile		Arg	Gln	Lys	Arg	Ile	Leu	Val	Asn	Leu	Ser	Met	
					75		•			80					85		
															AAG		403
	Va1	Glu	Asn	Lys	Leu	Val	Glu					Leu	Leu	Ser	Lys	Gly	
				90					95					100			
30	TTC	AGA	GGT	GCA	TCA	CCT	CAC	CGG	AAA	TCC	ACC	TAAA	AGCG	TAC	CAGG		450
•	Phe	Arg	Gly	Ala	Ser	Pro	His	Arg	Lys	Ser	Thr						
			105					110									
	ATGI	'AATC	CC A	GTGG	TGGA	IA A	CATI	`AAAC	AC	ACTI	TGA	GTAG	;				493
35																	
	(2)	INFO	RMAT	MOT	FOR	SEO	TD N	<u>ا</u> ۱۸۰۰	in•								

35

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 2044

,		(B) TYPE:	Nucleic aci	.d			
		(C) STRAND	EDNESS: Dou	ble		•	
		(D) TOPOLO	GY: Linear				
	(ii)	) SEQUENCE KI	ND: cDNA to	mRNA	•		
5					•		
	(vi)	ORIGINAL SO	URCE:				
		(A) ORGANI	SM: <i>Homo sa</i>	piens			
		(B) CELL K	IND: Epider	moid carci	noma		
		(C) CELL L	INE: KB				
10		(D) CLONE	NAME: HP104	28			
	(ix)	SEQUENCE CH	ARACTERISTI	CS:			
		(A) CHARAC	TERIZATION	CODE: CDS			
		(B) EXISTE	NCE POSITIO	N: 288 1	385		
15		(C) CHARAC	TERIZATION	METHOD: E			
	(xi)	SEQUENCE DE	SCRIPTION:	SEQ ID NO:	50:		
•		CTGGAGCTCC					60
20		GGGAAGGGTC .					120
		ATGGTTCCGA .					180
		COCTGTGGTCT .					240
	160016000	CGCTGACTCA	3GAGC1CCGG	16C1GCAGCC		Gly Arg	296
25					net 1	GIY AIR	
2.7	TGG GCC CT	C GAT GTG GC	ን ጥጥጥ ጥጥር ጥረ	36 AAG GCG	_	CTG GGG	344
		u Asp Val Ala					344
	5		10	•		Dea oxy	
	CTG GTG CT	T CTC TAC TAC				TAC AAC	392
30		u Leu Tyr Ty					
	20	2:		30		35	
		G ACA AAG AG		•	TTC ATG ACG		440
		u Thr Lys Se					
	•	40		45		50	
35	CAC CTG GC	C GTG ATC TT	CTC TTC TO	CC GCC CTG	TCC AGG GCG	CTG GTT	488
	His Leu Al	a Val Ile Pho	Leu Phe Se	er Ala Leu	Ser Arg Ala	Leu Val	
		55	(	50	65		
	CAG TGC TC	C AGC CAC AGG	GCC CGT G	rg gtg ctg	AGC TGG GCC	GAC TAC	536

	Gln	Cys	Ser	Şer	His	Arg	Ala	Arg	Val	Val	Leu	Ser	Trp	Ala	Asp	Tyr	,
			70					75					80				
	CTC	AGA	AGA	GTG	GCT	ccc	ACA	GCT	CTG	GCG	ACG	GCG	CTT	GAC	GTG	GGC	584
	Leu	Arg	Arg	Val	Ala	Pro	Thr	Ala	Leu	Ala	Thr	Ala	Leu	Asp	Val	Gly	
5		85					90					95					
	TTG	TCC	AAC	TGG	AGC	TTC	CTG	TAT	GTC	ACC	GTC	TCG	CTG	TAC	ACA	ATG	632
	Leu	Ser	Asn	Trp	Ser	Phe	Leu	Tyr	Val	Thr	Val	Ser	Leu	Tyr	Thr	Met	
	100					105					110	٠				115	
	ACC	AAA	TCC	TCA	GCT	GTC	CTC	TTC	ATC	TTG	ATC	TTC	TCT	CTG	ATC	TTC	680
10	Thr	Lys	Ser	Ser	Ala	Val	Leu	Phe	Ile	Leu	Ile	Phe	Ser	Leu	Ile	Phe	
					120					125					130		
•	AAG	CTG	GAG	GAG	CTG	CGC	GCG	GCA	CTG	GTC	CTG	GTG	GTC	CTC	CTC	ATC	728
	Lys	Leu	Glu	Glu	Leu	Arg	Ala	Ala	Leu	Val	Leu	Val	Val	Leu	Leu	Ile	
				135					140					145			
15	GCC	GGG	GGT	CTC	TTC	ATG	TTC	ACC	TAC	AAG	TCC	ACA	CAG	TTC	AAC	GTG	776
	Ala	Gly	Gly	Leu	Phe	Met	Phe	Thr	Tyr	Lys	Ser	Thr	Gln	Phe	Asn	Val	•
			150					155					160				
	GAG	GGC	TTC	GCC	TTG	GTG	CTG	GGG	GCC	TCG	TTC	ATC	GGT	GGC	ATT	CGC	824
	G1u	Gly	Phe	Ala	Leu	Val	Leu	Gly	Ala	Ser	Phe	Ile	Gly	Gly	Ile	Arg	
20		165					170					175					
	TGG	ACC	CTC	ACC	CAG	ATG	CTC	CTG	CAG	AAG	GCT	GAA	CTC	GGC	CTC	CAG	872
	Trp	Thr	Leu	Thr	Gln	Met	Leu	Leu	Gln	Lys	Ala	Glu	Leu	Gly	Leu	Gln	
	180					185					190					195	
	AAT	CCC	ATC	GAC	ACC	ATG	TTC	CAC	CTG	CAG	CCA	CTC	ATG	TTC	CTG	GGG	920
25	Asn	Pro	Ile	Asp	Thr	Met	Phe	His	Leu	Gln	Pro	Leu	Met	Phe	Leu	Gly	
		•			200					205					210		
			CCT														968
	Leu	Phe	Pro	Leu	Phe	Ala	Val	Phe	Glu	Gly	Leu	His	Leu	Ser	Thr	Ser	
		•		215					220					225			
30	GAG	AAA	ATC	TTC	CGT	TTC	CAG	GAC	ACA	GGG	CTG	CTC	CTG	CGG	GTA	CTT	1016
•	G1u	Lys	Ile	Phe	Arg	Phe	G1n	Asp	Thr	Gly	Leu	Leu	Leu	Arg	Val	Leu	
			230					235					240				
	GGG	AGC	CTC	TTC	CTT	GGC	GGG	ATT	CTC	GCC	TTT	GGT	TTG	GGC	TTC	TCT	1064
	G1y	Ser	Leu	Phe	Leu	Gly	Gly	Ile	Leu	Ala	Phe	Gly	Leu	Gly	Phe	Ser	
35		245					250					255					
			CTC														1112
	Glu	Phe	Leu	Leu	Val	Ser	Arg	Thr	Ser	Ser	Leu	Thr	Leu	Ser	Ile	Ala	
	260					265					270					275	

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	GGC	ATT	TTT	AAG	GAA	GTC	TGC	ACT	TTG	CTG	TTG	GCA	GCT	CAT	CTG	CTG	1160
	Gly	Ile	Phe	Lys	Glu	Val	Cys	Thr	Leu	Leu	Leu	Ala	Ala	His	Leu	Leu	
					280					285					290		
	GGC	GAT	CAG	ATC	AGC	CTC	CTG	AAC	TGG	CTG	GGÇ	TTC	GCC	CTC	TGC	CTC	1208
5	Gly	Asp	Gln	Ile	Ser	Leu	Leu	Asn	Trp	Leu	G1y	Phe	Ala	Leu	Cys	Leu	
				295					300					305			
	TCG	GGA	ATA	TCC	CTC	CAC	GTT	GCC	CTC	AAA	GCC	CTG	CAT	TCC	AGA	GGT	1256
	Ser	Gly	Ile	Ser	Leu	His	Val	Ala	Leu	Lys	Ala	Leu	His	Ser	Arg	Gly	
			310					315					320				
10	GAT	GGT	GGC	CCC	AAG	GCC	TTG	AAG	GGG	CTG	GGC	TCC	AGC	CCC	GAC	CTG	1304
	Asp	Gly	Gly	Pro	Lys	Ala	Leu	Lys	Gly	Leu	Gly	Ser	Ser	Pro	Asp	Leu	
		325					330					335					
	GAG	CTG	CTG	CTC	CGG	AGC	AGC	CAG	CGG	GAG	GAA	GGT	GAC	AAT	GAG	GAG	1352
	Glu	Leu	Leu	Leu	Arg	Ser	Ser	Gln	Arg	Glu	Glu	Gly	Asp	Asn	Glu	Glu	
15	340					345					350					355	
	GAG	GAG	TAC	TTT	GTG	GCC	CAG	GGG	CAG	CAG	TGAC	CAGC	CA G	GGCA	TAAA		1400
	Glu	G1u	Tyr	Phe	Val	Ala	Gln	Gly	Gln	Gln						•	
					360					365							
	GGCT	TAGA	AG (	CAGGC	CACT	c cc	CAGC	CTGC	TGC	CAGC	ACT	CACI	GTGC	TC A	AGCC	GCCAG	1460
20																GCCAG	1520
	GGAC	TCAT	'GA (	TTTT	GCCC	C TC	CCTI	CAGA	GCC	TGGI	CAC	ACAA	.GGGG	CG A	GCAC	CAGGC	1580
																.GGGGA	1640
	GTGG	GCTG	GT 1	CTTC	CCAC	C AC	TTCC	CAGG	CTC	TGAC	AGC	CGAG	ACTO	AT T	TCCA	AGGCA	1700
	CAGC	AGCI	TT C	TAAA	LGGGA	C TG	AGTT	TGGA	CTG	GGTT	TTG	GACC	TCCA	.GG G	GCTG	GAGCT	1760
25	TCAT	CACC	TG G	GCAG	TGTC	T TI	TCTC	AGAG	AGC	AGGT	TTC	ATTT	TAGT	TT G	GAAA	TAAAT	1820
																ACAGT	1880
	GTGG	GCCI	'GG C	CTCI	CCTI	T CC	TTTC	CCTG	CCT	GGAG	CCT	TCTT	CAAA	TG T	CTGG	TCTTA	1940
														cc c	CAGT	GGGGC	2000
• •	CCCA	CTGC	AC C	TGCI	'GGCA	G GA	AATA	AATG	AAT	GTTT	ACT	GAGT					2044
30																	

(2) INFORMATION FOR SEQ ID NO: 51:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1043

35 (B) TYPE: Nucleic acid

(C) STRANDEDNESS: Double

(D) TOPOLOGY: Linear

(ii) SEQUENCE KIND: cDNA to mRNA

WO 98/55508

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo sapiens

				(B)	CEL	L KI	ND:	Ston	ach	cano	er						
				(D)	CLC	NE N	IAME:	HP1	.0429	•							
5																	
		(	ix)	SEQU	ENCE	CHA	RACT	ERIS	TICS	:							
				(A)	СНА	RACI	ERIZ	ATIO	N CO	DE:	CDS						
				(B)	EXI	STEN	CE P	OSIT	'ION:	157	8	37					
				(C)	СНА	RACT	ERIZ	AŢIO	N ME	THOD	: E						
10															•		
		(	xi)	SEQU	ENCE	DES	CRIP	TION	: SE	Q ID	NO:	51:					
	ATT	AGCA	TAA	CCCT	TCCT	CA G	GAAG	AGTG	A GA	TTTT	ATAT	TTG	ACAA	TAA	AGTG	TTAGAC	60
	TCC	ATTT	CTA	AATA	CCAG	AC T	TCAA	AAGA	AA T	GGTT	CAAA	AGT	GTTA	AAT	GAAG	ATATTO	120
15	CTT	TTTT	TGT	CCTA	GAGA	AC T	TATT	TTCC	T GT	GAAA	ATG	CCT	ACC	ACA	AAG	AAG	174
											Met	Pro	Thr	Thr	Lys	Lys	
											1				5		
				TTC													222
••	Thr	Leu	Met	Phe	Leu	Ser	Ser	Phe	Phe	Thr	Ser	Leu	Gly	Ser	Phe	Ile	
20				10					15					20			
				TCT													270
	Val	116		Ser	Ile	Leu	Gly			Ala	Trp	Ile		Ser	Thr	Ile	
	C0#	0 mm	25		mam	0.00	ma.	30					35		m		
25				GAC													318
2,5	Ara	40	Arg	Asp	ser	AIB		ASI	GIÀ	Ser	TIE		TIE	Thr	Tyr	GIÀ	
	C do do		CCT	GGG	CAC	ለርጥ	45 40T	C	C 4 4	mmc.	A C (T)	50	004	CTT	CCA	C 4 4	366
				Gly													300
	55			019	GIU	60	Der	GIU	GIU	Deu	65	1112		neu	VIG	70	
30		AAG	AAA	AAG	TTT		GTT	ТТА	GAG	АТА		ААТ	ΑΑT	тст	TCC		414
				Lys													
		•		_, -	75					80					85		
	AAA	ACT	CTG	CAT		GTG	ACT	ATC	CTG		CTG	GTC	CTG	AGT		ATC	462
	_			His													
35				90					95					100			
	ACG	TCG	CTG	CTG	AGC	TCT	GGG	TTT	ACC	TTC	TAC	AAC	AGC	ATC	AGC	AAC	510
	Thr	Ser	Leu	Leu	Ser	Ser	Gly	Phe	Thr	Phe	Tyr	Asn	Ser	Ile	Ser	Asn	
			105					110					115				

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							•		. •	TOT							
	CCT	TAC	CAG	ACA	TTC	CTG	GGG	CCG	ACG	GGG	GTG	TAC	ACC	TGG	AAC	GGG	558
	Pro	Tyr	Gln	Thr	Phe	Leu	Gly	Pro	Thr	Gly	Val	Tyr	Thr	Trp	Asn	Gly	
		120					125					130					
	CTC	GGT	GCA	TCC	TTC	GTT	TTT	GTG	ACC	ATG	ATA	CTG	TTT	GTG	GCG	AAC	606
5	Leu	Gly	Ala	Ser	Phe	Val	Phe	Val	Thr	Met	Ile	Leu	Phe	Val	Ala	Asn	
	135					140					145					150	
	ACG	CAG	TCC	AAC	CAA	CTC	TCC	GAA	GAG	TTG	TTC	CAA	ATG	CTT	TAC	CCG	654
	Thr	Gln	Ser	Asn	Gln	Leu	Ser	Glu	Glu	Leu	Phe	Gln	Met	Leu	Tyr	Pro	
					155					160					165		
10	GCA	ACC	ACC	AGT	AAA	GGA	ACG	ACC	CAC	AGT	TAC	GGA	TAC	TCG	TTC	TGG	702
	Ala	Thr	Thr	Ser	Lys	Gly	Thr	Thr	His	Ser	Tyr	Gly	Tyr	Ser	Phe	Trp	
•				170					175					180			
	CTC	ATA	CTG	CTC	GTC	ATT	CTT	CTA	AAT	ATA	GTC	ACT	GTA	ACC	ATC	ATC	750
	Leu	Ile	Leu	Leu	Val	Ile	Leu	Leu	Asn	Ile	Val	Thr	Val	Thr	Ile	Ile	
15			185					190					195				
	ATT	TTC	TAC	CAG	AAG	GCC	AGA	TAC	CAG	CGG	AAG	CAG	GAG	CAG	AGA	AAG	798
	Ile	Phe	Tyr	Gln	Lys	Ala	Arg	Tyr	Gln	Arg	Lys	Gln	Glu	Gln	Arg	Lys	
		200					205					210					
	CCA	ATG	GAA	TAT	GCT	CCA	AGG	GAC	GGA	TTA	TTA	TTC	TGAA	TTCT	CT 1	TCATC	850
20	Pro	Met	Glu	Tyr	Ala	Pro	Arg	Asp	Gly	Ile	Leu	Phe					
	215					220					225						
	TCAT	PTTTG	GC G	TTG	CATC	IT AT	GTAC	CATCA	GCC	CTGA	GTA	GTAA	CTGG	TT A	GCTI	CTCTG	910
	GACA	ATTC	AG C	ATG	CAAT	G TG	ACTG	TCAT	CTG	TGAC	AGC	ATTT	GTGT	TT C	ATGA	CACTG	970
	TGT	CTTC	I TA	GATO	CTGT	CA CI	CCTG	AAAA	TTI	TTCC	CAC	AAGG	TTGG	GG A	AATG	AATGG	1030
25	GAAA	ATGTO	GC I	'GG													1043
								•								•	
	(2)	INFO	RMAT	NOI	FOR	SEQ	ID N	10: 5	2:								

(i) SEQUENCE CHARACTERISTICS:

30 (A) LENGTH: 972

- (B) TYPE: Nucleic acid
- (C) STRANDEDNESS: Double
- (D) TOPOLOGY: Linear
- (ii) SEQUENCE KIND: cDNA to mRNA

(vi) ORIGINAL SOURCE:

- (A) ORGANISM: Homo sapiens
- (B) CELL KIND: Liver

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## (D) CLONE NAME: HP10432

(i	x)	SEO	UENCE	CHARA	CT	ERI	STI	CS:
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5

- (A) CHARACTERIZATION CODE: CDS
- (B) EXISTENCE POSITION: 29.. 418
  - (C) CHARACTERIZATION METHOD: E

## (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 52:

• •																		
10	AGACA	AGCG	GC (	GGGC(	GCAG(	GA C	GTGC										5	52
								ı		Ala .	Arg	Gly		Leu	Arg	Arg		
	ጥጥሶ ሰ	- m-C	000	CEC.	·	CTIC	CMC	600	1	moo.	0.00	000	5 	- CBC		- mcc	10	١٥
	TTG C																10	,0
15	Leu I	10	Arg	Leu	Leu	ARI	Leu 15	GIÀ	Leu	Trp	Leu	A18		Leu	Arg	ser		
13	GTG G		GGG	GAG	CAA	GCG		ccc	۸۵۵	ccc	ccc			ccc	ccc	AGC	14	A
	Val A																14	
	25	110	Gly	GIU	GIII	30	FLU	Gly	1111	nia	35	cys	SET	vrR	Gly	40		
	TCC T	ກວດ	AGC	GCG	GAC		GAC	AAG	TGC	ATC		TGC	GCG	ጥርጥ	TGC		19	16
20	Ser T																19	•
		Р			45	Dea	p	11, 0	0,0	50	мор	0,3	1110	001	55			
	GCG C	GA	CCG	CAC	• -	GAC	TTC	TGC	CTG		TGC	GCT	GCA	GCA			24	4
	Ala A	•																·
		•		60		•		•	65		•			70				
25	GCC C	cc	TTC	CGG	CTG	CTT	TGG	CCC	ATC	CTT	GGG	GGC	GCT	CTG	AGC	CTG	29	2 .
	Ala P	ro	Phe	Arg	Leu	Leu	Trp	Pro	Ile	Leu	Gly	Gly	Ala	Leu	Ser	Leu		
			75				•	80			•	•	85					
	ACC T	TC	GTG	CTG	GGG	CTG	CTT	TCT	GGC	TTT	TTG	GTC	TGG	AGA	CGA	TGC	34	0
	Thr P	he	Val	Leu	Gly	Leu	Leu	Ser	Gly	Phe	Leu	Val	Trp	Arg	Arg	Cys		
30		90				·	95		•			100	Ī	_				
	CGC A	.GG	AGA	GAG	AAG	TTC	ACC	ACC	ccc	ATA	GAG	GAG	ACC	GGC	GGA	GAG	38	8
	Arg A	rg	Arg	Glu	Lys	Phe	Thr	Thr	Pro	Ile	Glu	Glu	Thr	Gly	Gly	Glu		
	105					110					115					120		
	GGC T	'GC	CCA	GCT	GTG	GCG	CTG	ATC	CAG	TGAC	CA A	IGT (	GCCC	CCTG	CC A	CCGG	44	0
35	Gly C	ys	Pro	Ala	Val	Ala	Leu	Ile	Gln							•		
					125													
	GGCTC	GCC	CA C	TCAT	CATI	ra or	TCAT	CCAT	TCT	AGAG	GCCA	GTC'	rctg:	CCT (	CCCA	GACGCG	500	0
	GCGGG	AGC	CA A	GCTC	CTCC	A AC	CACA	AGGG	GGG	TGGG	GGG	CGG'	TGAA'	TCA (	CCTC'	TGAGGC	560	0

WO 98/55508	PCT/JP98/0244
W C 70/33300	PC 1/JP98/0244;

CTGGGCCCAG GGTTCAGGGG AACCTTCCAA GGTGTCTGGT TGCCCTGCCT CTGGCTCCAG

	AACAGAAAGG GAGCCTCACG CTGGCTCACA CAAAACAGCT GACACTGACT AAGGAACTGC	680
	AGCATTTGCA CAGGGGAGGG GGGTGCCCTC CTTCCTAGAG GCCCTGGGGG CCAGGCTGAC	740
	TTGGGGGGCA GACTTGACAC TAGGCCCCAC TCACTCAGAT GTCCTGAAAT TCCACCACGG	800
5	GGGTCACCCT GGGGGGTTAG GGACCTATTT TTAACACTAG GGGGCTGGCC CACTAGGAGG	860
	GCTGGCCCTA AGATACAGAC CCCCCAACT CCCCAAAGCG GGGAGGAGAT ATTTATTTTG	920
	GGGAGAGTTT GGAGGGGAGG GAGAATTTAT TAATAAAAGA ATCTTTAACT TT	972
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	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 695	
	(B) TYPE: Nucleic acid	
	(C) STRANDEDNESS: Double	
15	(D) TOPOLOGY: Linear	
	(ii) SEQUENCE KIND: cDNA to mRNA	
	(vi) ORIGINAL SOURCE:	
20	(A) ORGANISM: Homo sapiens	
20	(B) CELL KIND: Liver	
	(C) CELL LINE: (D) CLONE NAME: HP10433	
	(D) CLONE NAME: DF10455	
	(ix) SEQUENCE CHARACTERISTICS:	
25	(A) CHARACTERIZATION CODE: CDS	
	(B) EXISTENCE POSITION: 73 564	
	(C) CHARACTERIZATION METHOD: E	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 53:	
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	TGACACGGAA GC ATG CGA CGG CTG CTG ATC CCT CTG GCC CTG TGG CTG GGC	111
	Met Arg Arg Leu Leu Ile Pro Leu Ala Leu Trp Leu Gly	
	1 5 10	
35	GCG GTG GGC GTC GCC GAG CTC ACG GAA GCC CAG CGC CGG GGC	159
	Ala Val Gly Val Gly Val Ala Glu Leu Thr Glu Ala Gln Arg Gly	
	15 20 25	
	CTG CAG GTG GCC CTG GAG GAA TTT CAC AAG CAC CCG CCC GTG CAG TGG	207

	Leu	Gln	Val	Ala	Leu	Glu	Glu	Phe	His	Lys	His	Pro	Pro	Val	Gln	Trp		
	30					35					40					45		
	GCC	TTC	CAG	GAG	ACC	AGT	GTG	GAG	AGC	GCC	GTG	GAC	ACG	CCC	TTC	CCA		255
	Ala	Phe	G1n	Glu	Thr	Ser	Val	Glu	Ser	Ala	Val	Asp	Thr	Pro	Phe	Pro		
5					50					55					60			
	GĊT	GGA	ATA	TTT	GTG	AGG	CTG	GAA	TTT	AAG	CTG	CAG	CAG	ACA	AGC	TGC		303
	Ala	Gly	Ile	Phe	Val	Arg	Leu	Glu	Phe	Lys	Leu	Gln	Gln	Thr	Ser	Cys		
				65					70					75				
	CGG	AAG	AGG	GAC	TGG	AAG	AAA	CCC	GAG	TGC	AAA	GTC	AGG	CCC	AAT	GGG		351
10 .	Arg	Lys	Arg	Asp	Trp	Lys	Lys	Pro	Glu	Cys	Lys	Val	Arg	Pro	Asn	Gly		
			80					85					90					
	AGG	AAA	CGG	AAA	TGC	CTG	GCC	TGC	ATC	AAA	CTG	GGC	TCT	GAG	GAC	AAA		399
	Arg	Lys	Arg	Lys	Cys	Leu	Ala	Cys	Ile	Lys	Leu	Gly	Ser	Glu	Asp	Lys		
		95					100					105						
15	GTT	CTG	GGC	CGG	TTG	GTC	CAC	TGC	CCC	ATA	GAG	ACC	CAA	GT <sub>T</sub>	CTG	CGG		447
	Val	Leu	Gly	Arg	Leu	Val	His	Cys	Pro	Ile	Glu	Thr	Gln	Val	Leu	Arg		
	110					115					120					125		
													GTG					495
	Glu	Ala	Glu	Glu		Gln	Glu	Thr	Gln	Cys	Leu	Arg	Val	Gln		Ala		
20					130					135	·				140			
													TTC					543
	Gly	Glu	Asp		His	Ser	Phe	Tyr		Pro	Gly	Gln	Phe		Phe	Ser		
		000		145					150					155				500
2 5							TAAG	CCAG	CA (	TGAG	CTGC	G TO	GTGC	CTC				590
25	гав	ALA		Pro	Arg	ser												
	CACC		160	2000	- maan	DA 4.0	2040	1004		2000		000				-CCC-01		650
														IGA C	GACC	CCGT	L	650 695
	CIAI		JAG (	CAIC	MIM	AT AA	MGC	GCIC	, 100	CAGC	166	CICI	ıc					093
30																		
50	(2)	TNF	חמאר) אמים	אסדי	FOR	SEQ	TD N	JO • 5										
	(2)					HARA												
		(-	., 51	•		TH:												
						: Nu			:10									
35						NDEI				•								
-				•														
	(D) TOPOLOGY: Linear (ii) SEQUENCE KIND: cDNA to mRNA																	
				•														

		(7	vi) (	ORIG	INAL	sou	RCE:										
				(A)	ORG	ANIS	M: H	omo .	sapi	ens							
				(B)	CEL	L KI	ND:	Stom	ach (	cance	er		•				
				(D)	CLO	NE N	AME:	HP1	0480								
5																	
		(:	ix) S	EQUI	ENCE	CHAI	RACT	ERIS'	TICS	:							
				(A)	CHAI	RACT	ERIZA	ATIO	N COI	DE: (	CDS						
				(B)	EXI	STEN	CE P	OSIT:	ION:	80.	. 66	1					
				(C)	CHAI	RACT	ERIZA	ATIO	N ME'	CHOD	: E						
10									•							•	
		()	ki) S	EQUI	ENCE	DES	CRIP	rion	: SEC	J ID	NO:	54:					
												0.50		200	0m000	208000	s 60
																CTCG(	112
1 =	CCC	CGCGC	JCG (	افاتاتان	rcaac											TGC Cys	112
15							. 110 1	s wr	g cy:	_	, ге. 5	1 DI	a Cy.	5 62.	1 11.		
	CGC	TGG	ATC	CTG	ccc		_	CTA	CTC			ATC	GCC	TTC	GAC		160
															Asp		
	6			15					20					25	_		
20	ATC	GCG	CTG	GCC	GGC	CGC	GGC	TGG	TTG	CAG	TCT	AGC	GAC	CAC	GGC	CAG	208
	Ile	Ala	Leu	Ala	Gly	Arg	Gly	Trp	Leu	Gln	Ser	Ser	Asp	His	Gly	Gln	
			30					35					40				
	ACG	TCC	TCG	CTG	TGG	TGG	AAA	TGC	TCC	CAA	GAG	GGC	GGC	GGC	AGC	GGG	256
	Thr	Ser	Ser	Leu	Trp	Trp	Lys	Cys	Ser	Gln	Glu	Gly	Gly	G1y	Ser	Gly	
25		45					50					55					
															GGT		304
	Ser	Tyr	Glu	Glu	Gly	Cys	Gln	Ser	Leu	Met		Tyr	Ala	Trp	Gly		
	60					65					70			0.00	4 m.a	75 man	250
20															ATC		352
30	Ala	A18	Ala	Ala		Leu	Phe	Cys	GIY		116	116	Leu	ANT	Ile 90	Cys	
	mmc.	A TI C	CTC	mcc	80	mmo	000	CTC	መረ ጥ	85	ccc	CAC	A TO	ርጥጥ	GŤC	<b>ጥጥ</b> ር	400
															Val		400
	rne	116	Leu	95	FILE	FILE	MIG	Leu	100	GLY	110	<b>G111</b>	1166	105			
35	СТС	AGA	стс		GGA	ССТ	СТС	ርጥጥ		TTG	GCT	GCT	GTG		CAG	ATC	448
<b>.</b> .															Gln		
		0	110		-3	· - <b>/</b>		115					120				
	ATC	TCC		GTA	ATT	TAC	CCC	GTG	AAG	TAC	ACC	CAG	ACC	TTC	ACC	CTT	496

	Ile Ser Leu Val Ile Tyr Pro Val Lys Tyr Thr Gln Thr Phe Thr Leu	
	125 130 135	
	CAT GCC AAC CGT GCT GTC ACT TAC ATC TAT AAC TGG GCC TAC GGC TTT	544
	His Ala Asn Arg Ala Val Thr Tyr Ile Tyr Asn Trp Ala Tyr Gly Phe	
5	140 145 150 155	
	GGG TGG GCA GCC ACG ATT ATC CTG ATC GGC TGT GCC TTC TTC TGC	592
	Gly Trp Ala Ala Thr Ile Ile Leu Ile Gly Cys Ala Phe Phe Cys	
	160 165 170	
	TGC CTC CCC AAC TAC GAA GAT GAC CTT CTG GGC AAT GCC AAG CCC AGG	640
10	Cys Leu Pro Asn Tyr Glu Asp Asp Leu Leu Gly Asn Ala Lys Pro Arg	
	175 180 185	
	TAC TTC TAC ACA TCT GCC TA ACTTGGG AATGAATGTG GGAGAAAATC GCT	690
	Tyr Phe Tyr Thr Ser Ala	
	190	
15	GCTGCTGAGA TGGACTCCAG AAGAAGAAAC TGTTTCTCCA GGCGACTTTG AACCCATTTT	750
	TTGGCAGTGT TCATATTATT AAACTAGTCA AAAATGCTAA AATAATTTGG GAGAAAATAT	810
	TTTTTAAGTA GTGTTATAGT TTCATGTTTA TCTTTTATTA TGTTTTGTGA AGTTGTGTCT	870
	TTTCACTAAT TACCTATACT ATGCCAATAT TTCCTTATAT CTATCCATAA CATTTATACT	930
٠.	ACATTTGTAA GAGAATATGC ACGTGAAACT TAACACTTTA TAAGGTAAAA ATGAGGTTTC	990
20	CAAGATTTAA TAATCTGATC AAGTTCTTGT TATTTCCAAA TAGAATGGAC TTGGTCTGTT	1050
	AAGGGCTAAG GAGAAGAGGA AGATAAGGTT AAAAGTTGTT AATGACCAAA CATTCTAAAA	1110
	GAAATGCAAA AAAAAAGTTT ATTTTCAAGC CTTCGAACTA TTTAAGGAAA GCAAAATCAT	1170
	TTCCTAAATG CATATCATTT GTGAGAATTT CTCATTAATA TCCTGAATCA TTCATTTCAG	1230
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25	TGTTGCCATA GTTGGTAAGG CTTTCCTTTA AGTGTGAAAT ATTTAGATGA AATTTTCTCT	1350
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	TTTGTGTTTA TATGTTCAGA ACCAGAGTAG ACTGGATTGA AAGATGGACT GGGTCTAATT	1470
	TATCATGACT GATAGATCTG GTTAAGTTGT GTAGTAAAGC ATTAGGAGGG TCATTCTTGT	1530
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	ACATGCCACA GGAGAATTCG GGGATTTGAG TTTCTCTGAA TAGCATATAT ATGATGCATC	1770
	GGATAGGTCA TTATGATTTT TTACCATTTC GACTTACATA ATGAAAACCA ATTCATTTTA	1830
	AATATCAGAT TATTATTTTG TAAGTTGTGG AAAAAGCTAA TTGTAGTTTT CATTATGAAG	1890
35	TTTTCCCAAT AAACCACCTA TTCT	1914

WO 98/55508

## CLAIMS

- A protein comprising an amino acid sequence selected from the group consisting of the amino acid sequences of SEQ
   ID NOS: 1 to 18.
  - 2. A DNA encoding the protein according to claim 1.
- 3. A cDNA comprising a nucleotide sequence selected from the group consisting of the nucleotide sequences of SEQ ID NOS: 19 to 36.
  - 4. A cDNA according to claim 3, which comprises a nucleotide sequence selected from the group consisting of the nucleotide sequences of SEQ ID NOS: 37 to 54.
  - 5. An expression vector capable of in vitro translating the DNA according to any of claims 2 to 4 or expressing said DNA in an eukaryotic cell.

20

15

6. A transformed eukaryotic cell capable of expressing the DNA according to any of claims 2 to 4 to produce the protein according to claim 1.

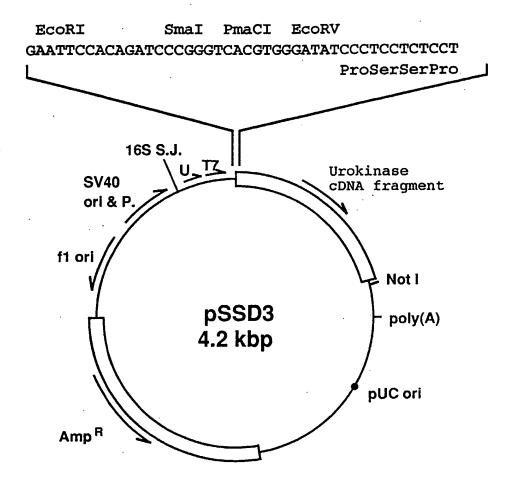


Fig.1

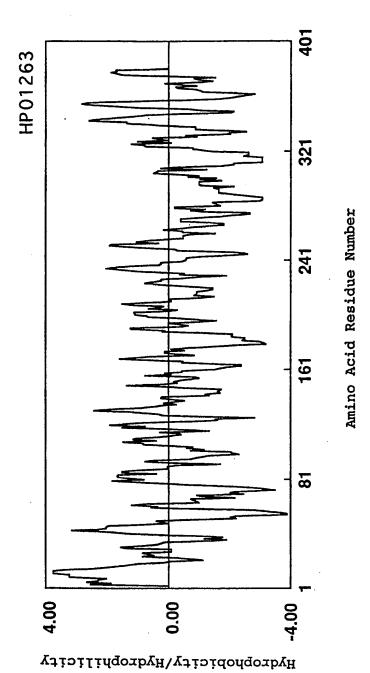


Fig.2

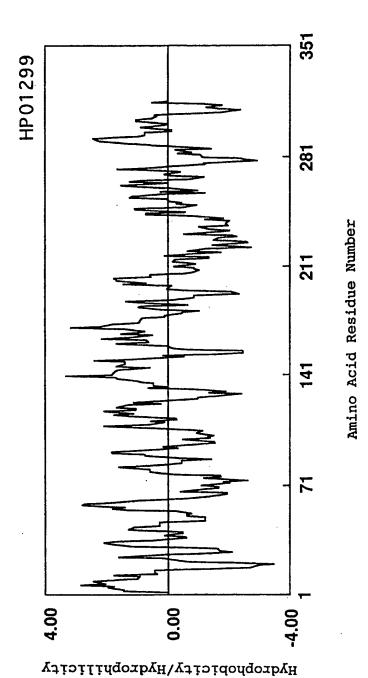


Fig.3

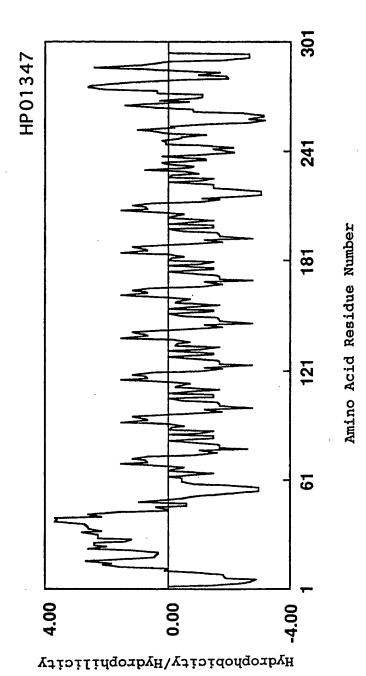


Fig.4

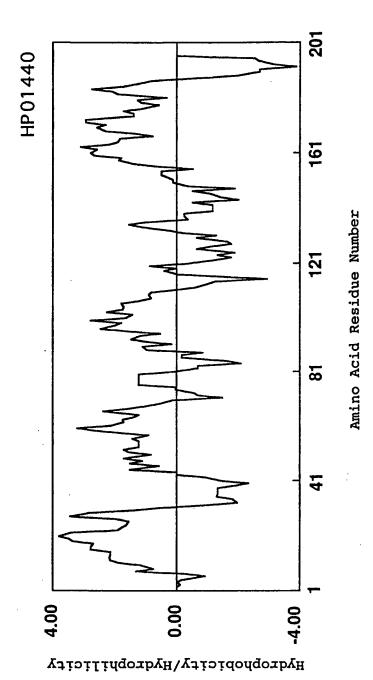


Fig.5

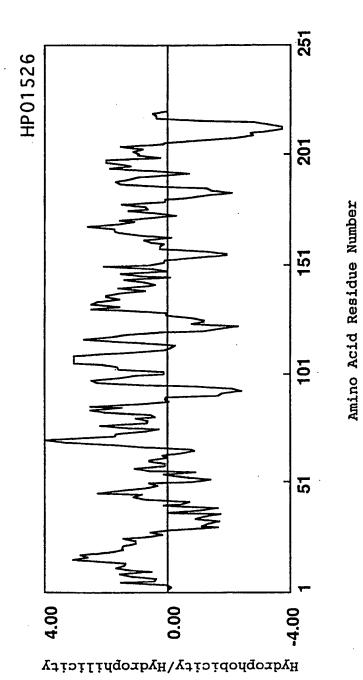


Fig.6

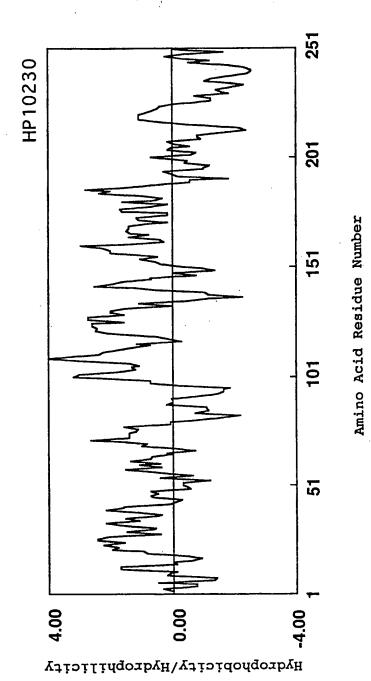


Fig.7

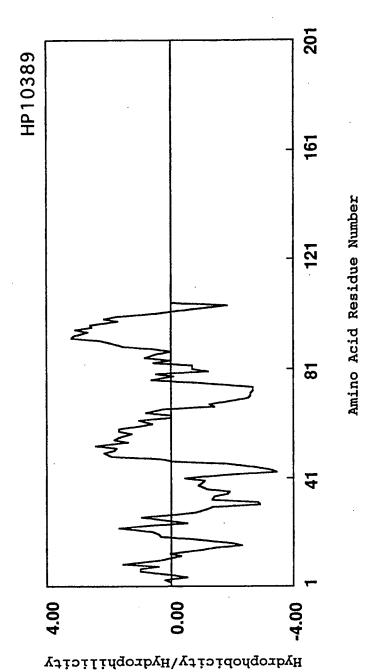


Fig.8

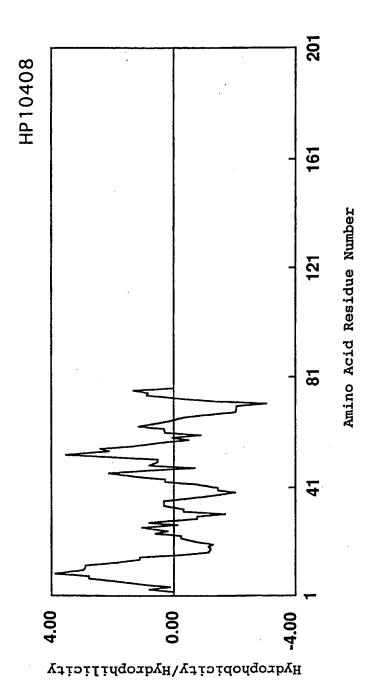


Fig.9

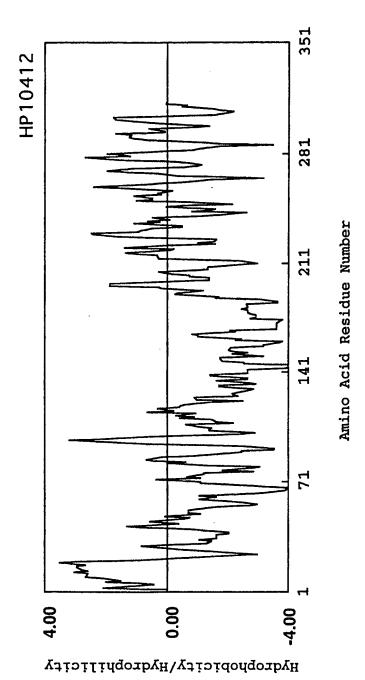


Fig.10

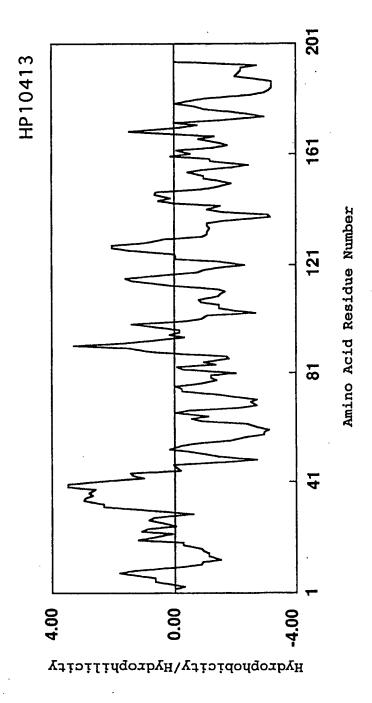


Fig.11

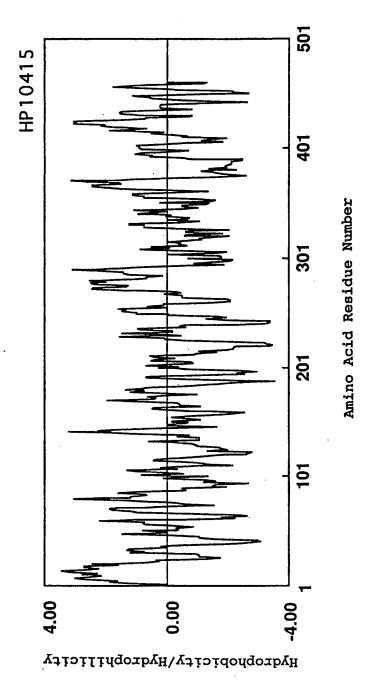


Fig.12

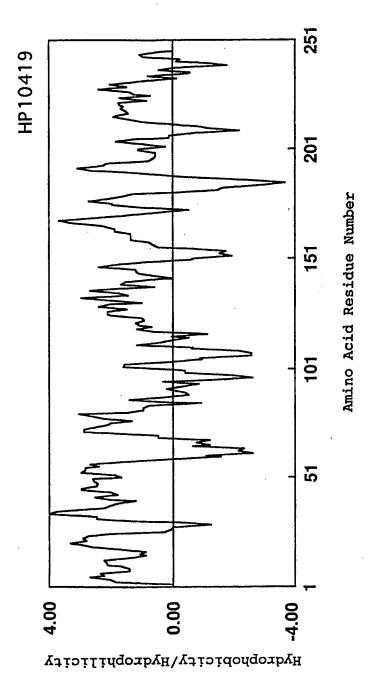


Fig.13

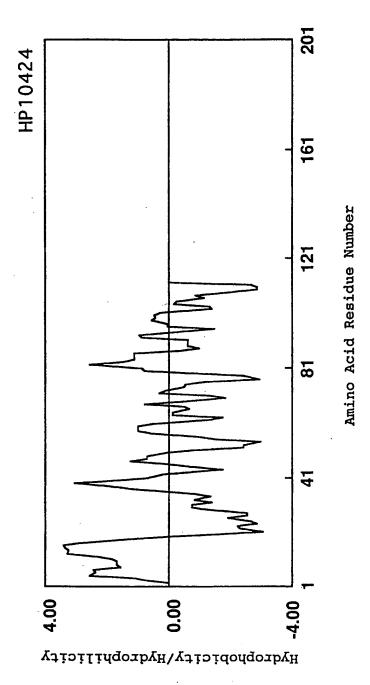


Fig.14

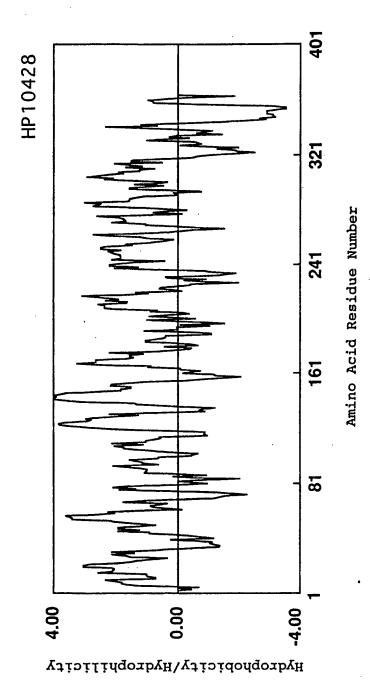
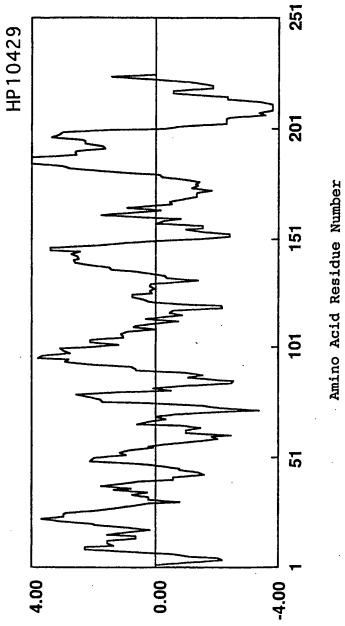


Fig.15



 ${\tt H} \lambda {\tt q} {\tt xobyoptc} {\tt r} {\tt f} \lambda {\tt h} {\tt q} {\tt xobyr} {\tt f} {\tt r} {\tt c} {\tt f} \lambda$ 

Fig.16

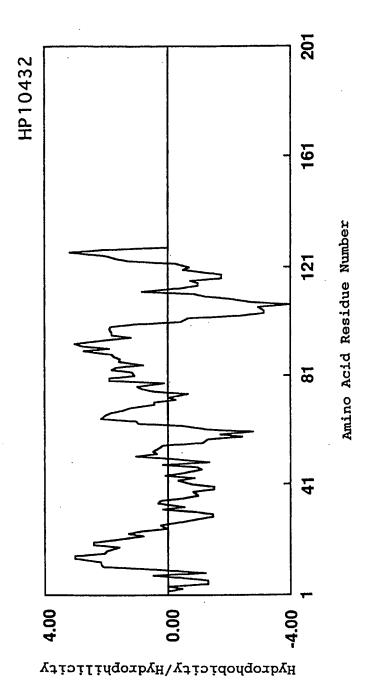
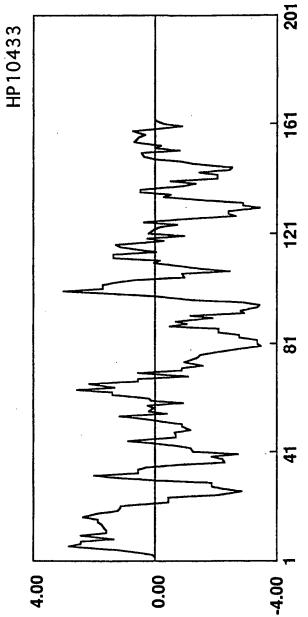


Fig.17



Amino Acid Residue Number

η Ηλακοδυορίατελ\Ηλακοδυίζιατελ

Fig.18

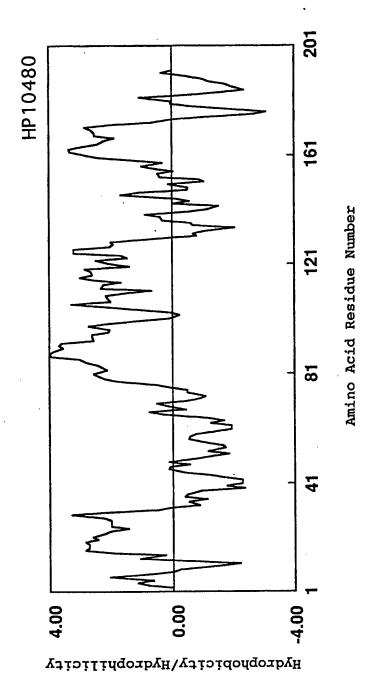


Fig.19